

to grow at 37 °C until an OD 600 of about 0.5 is reached. Take out an aliquot as T0 sample. Add 1 mM IPTG and allow to grow at 30 °C for 3 hours. Spin down the cells and store at -80 °C until purification. The determined cDNA and amino acid sequences for the P510S-C construct are shown in SEQ ID NO: 823 and 826, respectively.

- 5 The predicted third extracellular domain of P510S (P510S-E3; residues 328-676 of SEQ ID NO: 538) was expressed in *E. coli* as follows. The P510S fragment was amplified by PCR using the primers shown in SEQ ID NO: 830 and 831. The primer of SEQ ID NO: 830 is a sense primer with an NdeI site for use in ligating into pPDM. The primer of SEQ ID NO: 831 is an antisense primer with an added XhoI site
- 10 for use in ligating into pPDM. The resulting fragment was cloned to pPDM at the NdeI and XhoI sites. Clones were confirmed by sequencing. For protein expression, the clone was transformed into *E. coli* BL21 (DE3) CodonPlus-RIL competent cells. After induction, an OD600 of greater than 2.0 was achieved after 3 hours. Coomassie stained SDS-PAGE showed an over-expressed band at about 39 kD, and N-terminal sequencing
- 15 confirmed the N-terminal to be that of P510S-E3. Optimized culture conditions are as follows: dilute overnight culture/daytime culture (LB + kanamycin + chloramphenicol) into 2x YT (kanamycin and chloramphenicol) at a ratio of 25 ml culture to 1 liter 2x YT. Allow to grow at 37 °C until OD 600 equals 0.6. Take out an aliquot as T0 sample. Add 1 mM IPTG and allow to grow at 30 °C for 3 hours. Take out a T3
- 20 sample, spin down the cells and store at -80 °C until purification. The determined cDNA and amino acid sequences for the P501S-E3 construct are provided in SEQ ID NO: 824 and 827, respectively.

g) Expression of P775S in *E. Coli*

- The antigen P775P contains multiple open reading frames (ORF). The
- 25 third ORF, encoding the protein of SEQ ID NO: 483, has the best motif score. An expression fusion construct containing the *M. tuberculosis* antigen Ra12 (SEQ ID NO: 819) and P775P-ORF3 with an N-terminal 6x HisTag was prepared as follows. P775P-ORF3 was amplified using the sense PCR primers of SEQ ID NO: 832 and the antisense PCR primer of SEQ ID NO: 833. The PCR amplified fragment of P775P and

Ra12/pCRX1 were digested with the restriction enzymes EcoRI and XhoI. Vector and insert were ligated and then transformed into NovaBlue cells. Colonies were randomly screened for insert and then sequenced. A clone having the desired sequence was transformed into *E. coli* BL21 (DE3) CodonPlus-RIL competent cells. Two hours after induction, the cell density peaked at OD600 of approximately 1.8. Coomassie stained SDS-PAGE showed an over-expressed band at about 31 kD. Western blot using 6x HisTag antibody confirmed that the band was Ra12-P775P-ORF3. The determined cDNA and amino acid sequences for the fusion construct are provided in SEQ ID NO: 834 and 835, respectively.

H) Expression of a P703P His tag fusion protein in *E. coli*

The cDNA for the coding region of P703P was prepared by PCR using the primers of SEQ ID NO: 836 and 837. The PCR product was digested with EcoRI restriction enzyme, gel purified and cloned into a modified pET28 vector with a His tag in frame, which had been digested with Eco72I and EcoRI restriction enzymes. The correct construct was confirmed by DNA sequence analysis and then transformed into *E. coli* BL21 (DE3) pLys S expression host cells. The determined amino acid and cDNA sequences for the expressed recombinant P703P are provided in SEQ ID NO: 838 and 839, respectively.

I) Expression of a P705P His tag fusion protein in *E. coli*

The cDNA for the coding region of P705P was prepared by PCR using the primers of SEQ ID NO: 840 and 841. The PCR product was digested with EcoRI restriction enzyme, gel purified and cloned into a modified pET28 vector with a His tag in frame, which had been digested with Eco72I and EcoRI restriction enzymes. The correct construct was confirmed by DNA sequence analysis and then transformed into *E. coli* BL21 (DE3) pLys S and BL21 (DE3) CodonPlus expression host cells. The determined amino acid and cDNA sequences for the expressed recombinant P705P are provided in SEQ ID NO: 842 and 843, respectively.

J) Expression of a P711P His tag fusion protein in *E. coli*

The cDNA for the coding region of P711P was prepared by PCR using the primers of SEQ ID NO: 844 and 845. The PCR product was digested with EcoRI restriction enzyme, gel purified and cloned into a modified pET28 vector with a His tag in frame, which had been digested with Eco72I and EcoRI restriction enzymes. The correct construct was confirmed by DNA sequence analysis and then transformed into *E. coli* BL21 (DE3) pLys S and BL21 (DE3) CodonPlus expression host cells. The determined amino acid and cDNA sequences for the expressed recombinant P711P are provided in SEQ ID NO: 846 and 847, respectively.

K) Expression of P767P in *E. coli*

The full-length coding region of P767P (amino acids 2-374 of SEQ ID NO: 590) was amplified by PCR using the primers PDM-468 and PDM-469 (SEQ ID NO: 935 and 936, respectively). DNA amplification was performed using 10 µl 10X Pfu buffer, 1 µl 10 mM dNTPs, 2 µl each of the PCR primers at 10 µM concentration, 83 µl water, 1.5 µl Pfu DNA polymerase (Stratagene, La Jolla, CA) and 1 µl DNA at 100 ng/µl. Denaturation at 96°C was performed for 2 min, followed by 40 cycles of 96°C for 20 sec, 66°C for 15 sec and by 72°C for 2 min., and lastly by 1 cycle of 72°C for 4 min. The PCR product was digested with XhoI and cloned into a modified pET28 vector with a histidine tag in frame on the 5' end that was digested with Eco72I and XhoI. The construct was confirmed to be correct through sequence analysis and transformed into *E. coli* BL21 pLysS and BL21 CodonPlus RP cells. The cDNA coding region for the recombinant B767P protein is provided in SEQ ID NO: 938, with the corresponding amino acid sequence being provided in SEQ ID NO: 941. The full-length P767P did not express at high enough levels for detection or purification.

A truncated coding region of P767P (referred to as B767P-B; amino acids 47-374 of SEQ ID NO: 590) was amplified by PCR using the primers PDM-573 and PDM-469 (SEQ ID NO: 937 and 936, respectively) and the PCR conditions described above for full-length P767P. The PCR product was digested with XhoI and cloned into the modified pET28 vector that was digested with Eco72I and XhoI. The

construct was confirmed to be correct through sequence analysis and transformed into *E. coli* BL21 pLysS and BL21 CodonPlus RP cells. The protein was found to be expressed in the inclusion body pellet. The coding region for the expressed B767P-B protein is provided in SEQ ID NO: 939, with the corresponding amino acid sequence being provided in SEQ ID NO: 940.

EXAMPLE 18

PREPARATION AND CHARACTERIZATION OF ANTIBODIES AGAINST PROSTATE-SPECIFIC POLYPEPTIDES

a) Preparation and Characterization of Polyclonal Antibodies against P703P, P504S and P509S

Polyclonal antibodies against P703P, P504S and P509S were prepared as follows.

Each prostate tumor antigen expressed in an *E. coli* recombinant expression system was grown overnight in LB broth with the appropriate antibiotics at 37°C in a shaking incubator. The next morning, 10 ml of the overnight culture was added to 500 ml to 2x YT plus appropriate antibiotics in a 2L-baffled Erlenmeyer flask. When the Optical Density (at 560 nm) of the culture reached 0.4-0.6, the cells were induced with IPTG (1 mM). Four hours after induction with IPTG, the cells were harvested by centrifugation. The cells were then washed with phosphate buffered saline and centrifuged again. The supernatant was discarded and the cells were either frozen for future use or immediately processed. Twenty ml of lysis buffer was added to the cell pellets and vortexed. To break open the *E. coli* cells, this mixture was then run through the French Press at a pressure of 16,000 psi. The cells were then centrifuged again and the supernatant and pellet were checked by SDS-PAGE for the partitioning of the recombinant protein. For proteins that localized to the cell pellet, the pellet was resuspended in 10 mM Tris pH 8.0, 1% CHAPS and the inclusion body pellet was washed and centrifuged again. This procedure was repeated twice more. The washed

inclusion body pellet was solubilized with either 8 M urea or 6 M guanidine HCl containing 10 mM Tris pH 8.0 plus 10 mM imidazole. The solubilized protein was added to 5 ml of nickel-chelate resin (Qiagen) and incubated for 45 min to 1 hour at room temperature with continuous agitation. After incubation, the resin and protein mixture were poured through a disposable column and the flow through was collected. The column was then washed with 10-20 column volumes of the solubilization buffer. The antigen was then eluted from the column using 8M urea, 10 mM Tris pH 8.0 and 300 mM imidazole and collected in 3 ml fractions. A SDS-PAGE gel was run to determine which fractions to pool for further purification.

As a final purification step, a strong anion exchange resin such as HiPrepQ (Biorad) was equilibrated with the appropriate buffer and the pooled fractions from above were loaded onto the column. Each antigen was eluted off the column with a increasing salt gradient. Fractions were collected as the column was run and another SDS-PAGE gel was run to determine which fractions from the column to pool. The pooled fractions were dialyzed against 10 mM Tris pH 8.0. The proteins were then vialed after filtration through a 0.22 micron filter and the antigens were frozen until needed for immunization.

Four hundred micrograms of each prostate antigen was combined with 100 micrograms of muramyl dipeptide (MDP). Every four weeks rabbits were boosted with 100 micrograms mixed with an equal volume of Incomplete Freund's Adjuvant (IFA). Seven days following each boost, the animal was bled. Sera was generated by incubating the blood at 4°C for 12-4 hours followed by centrifugation.

Ninety-six well plates were coated with antigen by incubating with 50 microliters (typically 1 microgram) of recombinant protein at 4 °C for 20 hours. 250 microliters of BSA blocking buffer was added to the wells and incubated at room temperature for 2 hours. Plates were washed 6 times with PBS/0.01% Tween. Rabbit sera was diluted in PBS. Fifty microliters of diluted sera was added to each well and incubated at room temperature for 30 min. Plates were washed as described above before 50 microliters of goat anti-rabbit horse radish peroxidase (HRP) at a 1:10000 dilution was added and incubated at room temperature for 30 min. Plates were again

washed as described above and 100 microliters of TMB microwell peroxidase substrate was added to each well. Following a 15 min incubation in the dark at room temperature, the colorimetric reaction was stopped with 100 microliters of 1N H₂SO₄ and read immediately at 450 nm. All polyclonal antibodies showed immunoreactivity to the appropriate antigen.

b) Preparation and Characterization of Antibodies against P501S

A murine monoclonal antibody directed against the carboxy-terminus of the prostate-specific antigen P501S was prepared as follows.

A truncated fragment of P501S (amino acids 355-526 of SEQ ID NO: 113) was generated and cloned into the pET28b vector (Novagen) and expressed in *E. coli* as a thioredoxin fusion protein with a histidine tag. The trx-P501S fusion protein was purified by nickel chromatography, digested with thrombin to remove the trx fragment and further purified by an acid precipitation procedure followed by reverse phase HPLC.

Mice were immunized with truncated P501S protein. Serum bleeds from mice that potentially contained anti-P501S polyclonal sera were tested for P501S-specific reactivity using ELISA assays with purified P501S and trx-P501S proteins. Serum bleeds that appeared to react specifically with P501S were then screened for P501S reactivity by Western analysis. Mice that contained a P501S-specific antibody component were sacrificed and spleen cells were used to generate anti-P501S antibody producing hybridomas using standard techniques. Hybridoma supernatants were tested for P501S-specific reactivity initially by ELISA, and subsequently by FACS analysis of reactivity with P501S transduced cells. Based on these results, a monoclonal hybridoma referred to as 10E3 was chosen for further subcloning. A number of subclones were generated, tested for specific reactivity to P501S using ELISA and typed for IgG isotype. The results of this analysis are shown below in Table V. Of the 16 subclones tested, the monoclonal antibody 10E3-G4-D3 was selected for further study.

Table V

Isotype analysis of murine anti-P501S monoclonal antibodies

Hybridoma clone	Isotype	Estimated [Ig] in supernatant ($\mu\text{g/ml}$)
4D11	IgG1	14.6
1G1	IgG1	0.6
4F6	IgG1	72
4H5	IgG1	13.8
4H5-E12	IgG1	10.7
4H5-EH2	IgG1	9.2
4H5-H2-A10	IgG1	10
4H5-H2-A3	IgG1	12.8
4H5-H2-A10-G6	IgG1	13.6
4H5-H2-B11	IgG1	12.3
10E3	IgG2a	3.4
10E3-D4	IgG2a	3.8
10E3-D4-G3	IgG2a	9.5
10E3-D4-G6	IgG2a	10.4
10E3-E7	IgG2a	6.5
8H12	IgG2a	0.6

- 5 The specificity of 10E3-G4-D3 for P501S was examined by FACS analysis. Specifically, cells were fixed (2% formaldehyde, 10 minutes), permeabilized (0.1% saponin, 10 minutes) and stained with 10E3-G4-D3 at 0.5 – 1 $\mu\text{g/ml}$, followed by incubation with a secondary, FITC-conjugated goat anti-mouse Ig antibody (Pharmingen, San Diego, CA). Cells were then analyzed for FITC fluorescence using
- 10 an Excalibur fluorescence activated cell sorter. For FACS analysis of transduced cells, B-LCL were retrovirally transduced with P501S. For analysis of infected cells, B-LCL were infected with a vaccinia vector that expresses P501S. To demonstrate specificity in these assays, B-LCL transduced with a different antigen (P703P) and uninfected B-LCL vectors were utilized. 10E3-G4-D3 was shown to bind with P501S-transduced B-
- 15 LCL and also with P501S-infected B-LCL, but not with either uninfected cells or P703P-transduced cells.

To determine whether the epitope recognized by 10E3-G4-D3 was found on the surface or in an intracellular compartment of cells, B-LCL were transduced with P501S or HLA-B8 as a control antigen and either fixed and permeabilized as described

above or directly stained with 10E3-G4-D3 and analyzed as above. Specific recognition of P501S by 10E3-G4-D3 was found to require permeabilization, suggesting that the epitope recognized by this antibody is intracellular.

The reactivity of 10E3-G4-D3 with the three prostate tumor cell lines Lncap, PC-3 and DU-145, which are known to express high, medium and very low levels of P501S, respectively, was examined by permeabilizing the cells and treating them as described above. Higher reactivity of 10E3-G4-D3 was seen with Lncap than with PC-3, which in turn showed higher reactivity than DU-145. These results are in agreement with the real time PCR and demonstrate that the antibody specifically recognizes P501S in these tumor cell lines and that the epitope recognized in prostate tumor cell lines is also intracellular.

Specificity of 10E3-G4-D3 for P501S was also demonstrated by Western blot analysis. Lysates from the prostate tumor cell lines Lncap, DU-145 and PC-3, from P501S-transiently transfected HEK293 cells, and from non-transfected HEK293 cells were generated. Western blot analysis of these lysates with 10E3-G4-D3 revealed a 46 kDa immunoreactive band in Lncap, PC-3 and P501S-transfected HEK cells, but not in DU-145 cells or non-transfected HEK293 cells. P501S mRNA expression is consistent with these results since semi-quantitative PCR analysis revealed that P501S mRNA is expressed in Lncap, to a lesser but detectable level in PC-3 and not at all in DU-145 cells. Bacterially expressed and purified recombinant P501S (referred to as P501SStr2) was recognized by 10E3-G4-D3 (24 kDa), as was full-length P501S that was transiently expressed in HEK293 cells using either the expression vector VR1012 or pCEP4. Although the predicted molecular weight of P501S is 60.5 kDa, both transfected and "native" P501S run at a slightly lower mobility due to its hydrophobic nature.

Immunohistochemical analysis was performed on prostate tumor and a panel of normal tissue sections (prostate, adrenal, breast, cervix, colon, duodenum, gall bladder, ileum, kidney, ovary, pancreas, parotid gland, skeletal muscle, spleen and testis). Tissue samples were fixed in formalin solution for 24 hours and embedded in paraffin before being sliced into 10 micron sections. Tissue sections were permeabilized and incubated with 10E3-G4-D3 antibody for 1 hr. HRP-labeled anti-

mouse followed by incubation with DAB chromogen was used to visualize P501S immunoreactivity. P501S was found to be highly expressed in both normal prostate and prostate tumor tissue but was not detected in any of the other tissues tested.

To identify the epitope recognized by 10E3-G4-D3, an epitope mapping approach was pursued. A series of 13 overlapping 20-21 mers (5 amino acid overlap; SEQ ID NO: 489-501) was synthesized that spanned the fragment of P501S used to generate 10E3-G4-D3. Flat bottom 96 well microtiter plates were coated with either the peptides or the P501S fragment used to immunize mice, at 1 microgram/ml for 2 hours at 37 °C. Wells were then aspirated and blocked with phosphate buffered saline containing 1% (w/v) BSA for 2 hours at room temperature, and subsequently washed in PBS containing 0.1% Tween 20 (PBST). Purified antibody 10E3-G4-D3 was added at 2 fold dilutions (1000 ng – 16 ng) in PBST and incubated for 30 minutes at room temperature. This was followed by washing 6 times with PBST and subsequently incubating with HRP-conjugated donkey anti-mouse IgG (H+L) Affinipure F(ab') fragment (Jackson ImmunoResearch, West Grove, PA) at 1:20000 for 30 minutes. Plates were then washed and incubated for 15 minutes in tetramethyl benzidine. Reactions were stopped by the addition of 1N sulfuric acid and plates were read at 450 nm using an ELISA plate reader. As shown in Fig. 8, reactivity was seen with the peptide of SEQ ID NO: 496 (corresponding to amino acids 439-459 of P501S) and with the P501S fragment but not with the remaining peptides, demonstrating that the epitope recognized by 10E3-G4-D3 is localized to amino acids 439-459 of SEQ ID NO: 113.

In order to further evaluate the tissue specificity of P501S, multi-array immunohistochemical analysis was performed on approximately 4700 different human tissues encompassing all the major normal organs as well as neoplasias derived from these tissues. Sixty-five of these human tissue samples were of prostate origin. Tissue sections 0.6 mm in diameter were formalin-fixed and paraffin embedded. Samples were pretreated with HIER using 10 mM citrate buffer pH 6.0 and boiling for 10 min. Sections were stained with 10E3-G4-D3 and P501S immunoreactivity was visualized with HRP. All the 65 prostate tissues samples (5 normal, 55 untreated prostate tumors,

5 hormone refractory prostate tumors) were positive, showing distinct perinuclear staining. All other tissues examined were negative for P501S expression.

c) Preparation and Characterization of Antibodies against P503S

- 5 A fragment of P503S (amino acids 113-241 of SEQ ID NO: 114) was expressed and purified from bacteria essentially as described above for P501S and used to immunize both rabbits and mice. Mouse monoclonal antibodies were isolated using standard hybridoma technology as described above. Rabbit monoclonal antibodies were isolated using Selected Lymphocyte Antibody Method (SLAM) technology at
10 Immgenics Pharmaceuticals (Vancouver, BC, Canada). Table VI, below, lists the monoclonal antibodies that were developed against P503S.

Table VI

Antibody	Species
20D4	Rabbit
JA1	Rabbit
1A4	Mouse
1C3	Mouse
1C9	Mouse
1D12	Mouse
2A11	Mouse
2H9	Mouse
4H7	Mouse
8A8	Mouse
8D10	Mouse
9C12	Mouse
6D12	Mouse

15

The DNA sequences encoding the complementarity determining regions (CDRs) for the rabbit monoclonal antibodies 20D4 and JA1 were determined and are provided in SEQ ID NO: 502 and 503, respectively.

In order to better define the epitope binding region of each of the antibodies, a series of overlapping peptides were generated that span amino acids 109-213 of SEQ ID NO: 114. These peptides were used to epitope map the anti-P503S monoclonal antibodies by ELISA as follows. The recombinant fragment of P503S that was employed as the immunogen was used as a positive control. Ninety-six well microtiter plates were coated with either peptide or recombinant antigen at 20 ng/well overnight at 4 °C. Plates were aspirated and blocked with phosphate buffered saline containing 1% (w/v) BSA for 2 hours at room temperature then washed in PBS containing 0.1% Tween 20 (PBST). Purified rabbit monoclonal antibodies diluted in PBST were added to the wells and incubated for 30 min at room temperature. This was followed by washing 6 times with PBST and incubation with Protein-A HRP conjugate at a 1:2000 dilution for a further 30 min. Plates were washed six times in PBST and incubated with tetramethylbenzidine (TMB) substrate for a further 15 min. The reaction was stopped by the addition of 1N sulfuric acid and plates were read at 450 nm using at ELISA plate reader. ELISA with the mouse monoclonal antibodies was performed with supernatants from tissue culture run neat in the assay.

All of the antibodies bound to the recombinant P503S fragment, with the exception of the negative control SP2 supernatant. 20D4, JA1 and ID12 bound strictly to peptide #2101 (SEQ ID NO: 504), which corresponds to amino acids 151-169 of SEQ ID NO: 114. 1C3 bound to peptide #2102 (SEQ ID NO: 505), which corresponds to amino acids 165-184 of SEQ ID NO: 114. 9C12 bound to peptide #2099 (SEQ ID NO: 522), which corresponds to amino acids 120-139 of SEQ ID NO: 114. The other antibodies bind to regions that were not examined in these studies.

Subsequent to epitope mapping, the antibodies were tested by FACS analysis on a cell line that stably expressed P503S to confirm that the antibodies bind to cell surface epitopes. Cells stably transfected with a control plasmid were employed as a negative control. Cells were stained live with no fixative. 0.5 ug of anti-P503S monoclonal antibody was added and cells were incubated on ice for 30 min before being washed twice and incubated with a FITC-labelled goat anti-rabbit or mouse secondary antibody for 20 min. After being washed twice, cells were analyzed with an Excalibur

fluorescent activated cell sorter. The monoclonal antibodies 1C3, 1D12, 9C12, 20D4 and JA1, but not 8D3, were found to bind to a cell surface epitope of P503S.

In order to determine which tissues express P503S, immunohistochemical analysis was performed, essentially as described above, on a panel of normal tissues (prostate, adrenal, breast, cervix, colon, duodenum, gall bladder, ileum, kidney, ovary, pancreas, parotid gland, skeletal muscle, spleen and testis). HRP-labeled anti-mouse or anti-rabbit antibody followed by incubation with TMB was used to visualize P503S immunoreactivity. P503S was found to be highly expressed in prostate tissue, with lower levels of expression being observed in cervix, colon, ileum and kidney, and no expression being observed in adrenal, breast, duodenum, gall bladder, ovary, pancreas, parotid gland, skeletal muscle, spleen and testis.

Western blot analysis was used to characterize anti-P503S monoclonal antibody specificity. SDS-PAGE was performed on recombinant (rec) P503S expressed in and purified from bacteria and on lysates from HEK293 cells transfected with full length P503S. Protein was transferred to nitrocellulose and then Western blotted with each of the anti-P503S monoclonal antibodies (20D4, JA1, 1D12, 6D12 and 9C12) at an antibody concentration of 1 μ g/ml. Protein was detected using horse radish peroxidase (HRP) conjugated to either a goat anti-mouse monoclonal antibody or to protein A-sepharose. The monoclonal antibody 20D4 detected the appropriate molecular weight 14 kDa recombinant P503S (amino acids 113-241) and the 23.5 kDa species in the HEK293 cell lysates transfected with full length P503S. Other anti-P503S monoclonal antibodies displayed similar specificity by Western blot.

d) Preparation and Characterization of Antibodies against P703P

Rabbits were immunized with either a truncated (P703Ptr1; SEQ ID NO: 172) or full-length mature form (P703Pfl; SEQ ID NO: 523) of recombinant P703P protein was expressed in and purified from bacteria as described above. Affinity purified polyclonal antibody was generated using immunogen P703Pfl or P703Ptr1 attached to a solid support. Rabbit monoclonal antibodies were isolated using SLAM

technology at Immunogenics Pharmaceuticals. Table VII below lists both the polyclonal and monoclonal antibodies that were generated against P703P.

Table VII

Antibody	Immunogen	Species/type
Aff. Purif. P703P (truncated); #2594	P703PtrI	Rabbit polyclonal
Aff. Purif. P703P (full length); #9245	P703Pfl	Rabbit polyclonal
2D4	P703PtrI	Rabbit monoclonal
8H2	P703PtrI	Rabbit monoclonal
7H8	P703PtrI	Rabbit monoclonal

The DNA sequences encoding the complementarity determining regions (CDRs) for the rabbit monoclonal antibodies 8H2, 7H8 and 2D4 were determined and are provided in SEQ ID NO: 506-508, respectively.

Epitope mapping studies were performed as described above. Monoclonal antibodies 2D4 and 7H8 were found to specifically bind to the peptides of SEQ ID NO: 509 (corresponding to amino acids 145-159 of SEQ ID NO: 172) and SEQ ID NO: 510 (corresponding to amino acids 11-25 of SEQ ID NO: 172), respectively. The polyclonal antibody 2594 was found to bind to the peptides of SEQ ID NO: 511-514, with the polyclonal antibody 9427 binding to the peptides of SEQ ID NO: 515-517.

The specificity of the anti-P703P antibodies was determined by Western blot analysis as follows. SDS-PAGE was performed on (1) bacterially expressed recombinant antigen; (2) lysates of HEK293 cells and Ltk^{-/-} cells either untransfected or transfected with a plasmid expressing full length P703P; and (3) supernatant isolated from these cell cultures. Protein was transferred to nitrocellulose and then Western blotted using the anti-P703P polyclonal antibody #2594 at an antibody concentration of 1 µg/ml. Protein was detected using horse radish peroxidase (HRP) conjugated to an anti-rabbit antibody. A 35 kDa immunoreactive band could be observed with recombinant P703P. Recombinant P703P runs at a slightly higher molecular weight since it is epitope tagged. In lysates and supernatants from cells transfected with full length P703P, a 30 kDa band corresponding to P703P was observed. To assure

specificity, lysates from HEK293 cells stably transfected with a control plasmid were also tested and were negative for P703P expression. Other anti-P703P antibodies showed similar results.

Immunohistochemical studies were performed as described above, using anti-P703P monoclonal antibody. P703P was found to be expressed at high levels in normal prostate and prostate tumor tissue but was not detectable in all other tissues tested (breast tumor, lung tumor and normal kidney).

e) Preparation and Characterization of Antibodies against P504S

Full-length P504S (SEQ ID NO: 108) was expressed and purified from bacteria essentially as described above for P501S and employed to raise rabbit monoclonal antibodies using Selected Lymphocyte Antibody Method (SLAM) technology at Immgenics Pharmaceuticals (Vancouver, BC, Canada). The anti-P504S monoclonal antibody 13H4 was shown by Western blot to bind to both expressed recombinant P504S and to native P504S in tumor cells.

Immunohistochemical studies using 13H4 to assess P504S expression in various prostate tissues were performed as described above. A total of 104 cases, including 65 cases of radical prostatectomies with prostate cancer (PC), 26 cases of prostate biopsies and 13 cases of benign prostate hyperplasia (BPH), were stained with the anti-P504S monoclonal antibody 13H4. P504S showed strongly cytoplasmic granular staining in 64/65 (98.5%) of PCs in prostatectomies and 26/26 (100%) of PCs in prostatic biopsies. P504S was stained strongly and diffusely in carcinomas (4+ in 91.2% of cases of PC; 3+ in 5.5%; 2+ in 2.2% and 1+ in 1.1%) and high grade prostatic intraepithelial neoplasia (4+ in all cases). The expression of P504S did not vary with Gleason score. Only 17/91 (18.7%) of cases of NP/BPH around PC and 2/13 (15.4%) of BPH cases were focally (1+, no 2+ to 4+ in all cases) and weakly positive for P504S in large glands. Expression of P504S was not found in small atrophic glands, postatrophic hyperplasia, basal cell hyperplasia and transitional cell metaplasia in either biopsies or prostatectomies. P504S was thus found to be over-expressed in all Gleason scores of prostate cancer (98.5 to 100% of sensitivity) and exhibited only focal positivities in

large normal glands in 19/104 of cases (82.3% of specificity). These findings indicate that P504S may be usefully employed for the diagnosis of prostate cancer.

EXAMPLE 19

CHARACTERIZATION OF CELL SURFACE EXPRESSION AND CHROMOSOME LOCALIZATION OF THE PROSTATE-SPECIFIC ANTIGEN P501S

This example describes studies demonstrating that the prostate-specific antigen P501S is expressed on the surface of cells, together with studies to determine the probable chromosomal location of P501S.

The protein P501S (SEQ ID NO: 113) is predicted to have 11 transmembrane domains. Based on the discovery that the epitope recognized by the anti-P501S monoclonal antibody 10E3-G4-D3 (described above in Example 17) is intracellular, it was predicted that following transmembrane determinants would allow the prediction of extracellular domains of P501S. Fig. 9 is a schematic representation of the P501S protein showing the predicted location of the transmembrane domains and the intracellular epitope described in Example 17. Underlined sequence represents the predicted transmembrane domains, bold sequence represents the predicted extracellular domains, and italicized sequence represents the predicted intracellular domains. Sequence that is both bold and underlined represents sequence employed to generate polyclonal rabbit serum. The location of the transmembrane domains was predicted using HHMTOP as described by Tusnady and Simon (Principles Governing Amino Acid Composition of Integral Membrane Proteins: Applications to Topology Prediction, *J. Mol. Biol.* 283:489-506, 1998).

Based on Fig. 9, the P501S domain flanked by the transmembrane domains corresponding to amino acids 274-295 and 323-342 is predicted to be extracellular. The peptide of SEQ ID NO: 518 corresponds to amino acids 306-320 of P501S and lies in the predicted extracellular domain. The peptide of SEQ ID NO: 519, which is identical to the peptide of SEQ ID NO: 518 with the exception of the substitution of the histidine with an asparagine, was synthesized as described above. A

- Cys-Gly was added to the C-terminus of the peptide to facilitate conjugation to the carrier protein. Cleavage of the peptide from the solid support was carried out using the following cleavage mixture: trifluoroacetic acid:ethanediol:thioanisole:water:phenol (40:1:2:2:3). After cleaving for two hours, the peptide was precipitated in cold ether.
- 5 The peptide pellet was then dissolved in 10% v/v acetic acid and lyophilized prior to purification by C18 reverse phase hplc. A gradient of 5-60% acetonitrile (containing 0.05% TFA) in water (containing 0.05% TFA) was used to elute the peptide. The purity of the peptide was verified by hplc and mass spectrometry, and was determined to be >95%. The purified peptide was used to generate rabbit polyclonal antisera as described
- 10 above.

- Surface expression of P501S was examined by FACS analysis. Cells were stained with the polyclonal anti-P501S peptide serum at 10 µg/ml, washed, incubated with a secondary FITC-conjugated goat anti-rabbit Ig antibody (ICN), washed and analyzed for FITC fluorescence using an Excalibur fluorescence activated cell
- 15 sorter. For FACS analysis of transduced cells, B-LCL were retrovirally transduced with P501S. To demonstrate specificity in these assays, B-LCL transduced with an irrelevant antigen (P703P) or nontransduced were stained in parallel. For FACS analysis of prostate tumor cell lines, Lncap, PC-3 and DU-145 were utilized. Prostate tumor cell lines were dissociated from tissue culture plates using cell dissociation medium and
- 20 stained as above. All samples were treated with propidium iodide (PI) prior to FACS analysis, and data was obtained from PI-excluding (i.e., intact and non-permeabilized) cells. The rabbit polyclonal serum generated against the peptide of SEQ ID NO: 519 was shown to specifically recognize the surface of cells transduced to express P501S, demonstrating that the epitope recognized by the polyclonal serum is extracellular.

- 25 To determine biochemically if P501S is expressed on the cell surface, peripheral membranes from Lncap cells were isolated and subjected to Western blot analysis. Specifically, Lncap cells were lysed using a dounce homogenizer in 5 ml of homogenization buffer (250 mM sucrose, 10 mM HEPES, 1mM EDTA, pH 8.0, 1 complete protease inhibitor tablet (Boehringer Mannheim)). Lysate samples were spun
- 30 at 1000 g for 5 min at 4 °C. The supernatant was then spun at 8000g for 10 min at 4 °C.

Supernatant from the 8000g spin was recovered and subjected to a 100,000g spin for 30 min at 4 °C to recover peripheral membrane. Samples were then separated by SDS-PAGE and Western blotted with the mouse monoclonal antibody 10E3-G4-D3 (described above in Example 17) using conditions described above. Recombinant purified P501S, as well as HEK293 cells transfected with and over-expressing P501S were included as positive controls for P501S detection. LCL cell lysate was included as a negative control. P501S could be detected in Lucap total cell lysate, the 8000g (internal membrane) fraction and also in the 100,000g (plasma membrane) fraction. These results indicate that P501S is expressed at, and localizes to, the peripheral membrane.

To demonstrate that the rabbit polyclonal antiserum generated to the peptide of SEQ ID NO: 519 specifically recognizes this peptide as well as the corresponding native peptide of SEQ ID NO: 518, ELISA analyses were performed. For these analyses, flat-bottomed 96 well microtiter plates were coated with either the peptide of SEQ ID NO: 519, the longer peptide of SEQ ID NO: 520 that spans the entire predicted extracellular domain, the peptide of SEQ ID NO: 521 which represents the epitope recognized by the P501S-specific antibody 10E3-G4-D3, or a P501S fragment (corresponding to amino acids 355-526 of SEQ ID NO: 113) that does not include the immunizing peptide sequence, at 1 µg/ml for 2 hours at 37 °C. Wells were aspirated, blocked with phosphate buffered saline containing 1% (w/v) BSA for 2 hours at room temperature and subsequently washed in PBS containing 0.1% Tween 20 (PBST). Purified anti-P501S polyclonal rabbit serum was added at 2 fold dilutions (1000 ng - 125 ng) in PBST and incubated for 30 min at room temperature. This was followed by washing 6 times with PBST and incubating with HRP-conjugated goat anti-rabbit IgG (H+L) Affinipure F(ab') fragment at 1:20000 for 30 min. Plates were then washed and incubated for 15 min in tetramethyl benzidine. Reactions were stopped by the addition of 1N sulfuric acid and plates were read at 450 nm using an ELISA plate reader. As shown in Fig. 11, the anti-P501S polyclonal rabbit serum specifically recognized the peptide of SEQ ID NO: 519 used in the immunization as well as the longer peptide of

SEQ ID NO: 520, but did not recognize the irrelevant P501S-derived peptides and fragments.

In further studies, rabbits were immunized with peptides derived from the P501S sequence and predicted to be either extracellular or intracellular, as shown in Fig. 9. Polyclonal rabbit sera were isolated and polyclonal antibodies in the serum were purified, as described above. To determine specific reactivity with P501S, FACS analysis was employed, utilizing either B-LCL transduced with P501S or the irrelevant antigen P703P, of B-LCL infected with vaccinia virus-expressing P501S. For surface expression, dead and non-intact cells were excluded from the analysis as described above. For intracellular staining, cells were fixed and permeabilized as described above. Rabbit polyclonal serum generated against the peptide of SEQ ID NO: 548, which corresponds to amino acids 181-198 of P501S, was found to recognize a surface epitope of P501S. Rabbit polyclonal serum generated against the peptide SEQ ID NO: 551, which corresponds to amino acids 543-553 of P501S, was found to recognize an epitope that was either potentially extracellular or intracellular since in different experiments intact or permeabilized cells were recognized by the polyclonal sera. Based on similar deductive reasoning, the sequences of SEQ ID NO: 541-547, 549 and 550, which correspond to amino acids 109-122, 539-553, 509-520, 37-54, 342-359, 295-323, 217-274, 143-160 and 75-88, respectively, of P501S, can be considered to be potential surface epitopes of P501S recognized by antibodies.

In further studies, mouse monoclonal antibodies were raised against amino acids 296 to 322 to P501S, which are predicted to be in an extracellular domain. A/J mice were immunized with P501S/adenovirus, followed by subsequent boosts with an *E. coli* recombinant protein, referred to as P501N, that contains amino acids 296 to 322 of P501S, and with peptide 296-322 (SEQ ID NO: 898) coupled with KLH. The mice were subsequently used for splenic B cell fusions to generate anti-peptide hybridomas. The resulting 3 clones, referred to as 4F4 (IgG1,kappa), 4G5 (IgG2a,kappa) and 9B9 (IgG1,kappa), were grown for antibody production. The 4G5 mAb was purified by passing the supernatant over a Protein A-sepharose column,

followed by antibody elution using 0.2M glycine, pH 2.3. Purified antibody was neutralized by the addition of 1M Tris, pH 8, and buffer exchanged into PBS.

For ELISA analysis, 96 well plates were coated with P501S peptide 296-322 (referred to as P501-long), an irrelevant P775 peptide, P501S-N, P501TR2, P501S-long-KLH, P501S peptide 306-319 (referred to as P501-short)-KLH, or the irrelevant peptide 2073-KLH, all at a concentration of 2 ug/ml and allowed to incubate for 60 minutes at 37 °C. After coating, plates were washed 5X with PBS + 0.1% Tween and then blocked with PBS, 0.5% BSA, 0.4% Tween20 for 2 hours at room temperature. Following the addition of supernatants or purified mAb, the plates were incubated for 60 minutes at room temperature. Plates were washed as above and donkey anti-mouse IgHRP-linked secondary antibody was added and incubated for 30 minutes at room temperature, followed by a final washing as above. TMB peroxidase substrate was added and incubated 15 minutes at room temperature in the dark. The reaction was stopped by the addition of 1N H₂SO₄ and the OD was read at 450 nM. All three hybrid clones secreted mAb that recognized peptide 296-322 and the recombinant protein P501N.

For FACS analysis, HEK293 cells were transiently transfected with a P501S/VR1012 expression constructs using Fugene 6 reagent. After 2 days of culture, cells were harvested and washed, then incubated with purified 4C5 mAb for 30 minutes on ice. After several washes in PBS, 0.5% BSA, 0.01% azide, goat anti-mouse Ig-FITC was added to the cells and incubated for 30 minutes on ice. Cells were washed and resuspended in wash buffer including 1% propidium iodide and subjected to FACS analysis. The FACS analysis confirmed that amino acids 296-322 of P501S are in an extracellular domain and are cell surface expressed.

The chromosomal location of P501S was determined using the GeneBridge 4 Radiation Hybrid panel (Research Genetics). The PCR primers of SEQ ID NO: 528 and 529 were employed in PCR with DNA pools from the hybrid panel according to the manufacturer's directions. After 38 cycles of amplification, the reaction products were separated on a 1.2% agarose gel, and the results were analyzed through the Whitehead Institute/MIT Center for Genome Research web server

- (<http://www-genome.wi.mit.edu/cgi-bin/config/rhmapper.pl>) to determine the probable chromosomal location. Using this approach, P501S was mapped to the long arm of chromosome 1 at WI-9641 between q32 and q42. This region of chromosome 1 has been linked to prostate cancer susceptibility in hereditary prostate cancer (Smith *et al.* 5 *Science* 274:1371-1374, 1996 and Berthon *et al. Am. J. Hum. Genet.* 62:1416-1424, 1998). These results suggest that P501S may play a role in prostate cancer malignancy.

EXAMPLE 20

REGULATION OF EXPRESSION OF THE PROSTATE-SPECIFIC ANTIGEN P501S

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Steroid (androgen) hormone modulation is a common treatment modality in prostate cancer. The expression of a number of prostate tissue-specific antigens have previously been demonstrated to respond to androgen. The responsiveness of the prostate-specific antigen P501S to androgen treatment was examined in a tissue culture 15 system as follows.

Cells from the prostate tumor cell line LNCaP were plated at 1.5×10^5 cells/T75 flask (for RNA isolation) or 3×10^5 cells/well of a 6-well plate (for FACS analysis) and grown overnight in RPMI 1640 media containing 10% charcoal-stripped fetal calf serum (BRL Life Technologies, Gaithersburg, MD). Cell culture was 20 continued for an additional 72 hours in RPMI 1640 media containing 10% charcoal-stripped fetal calf serum, with 1 nM of the synthetic androgen Methyltrienolone (R1881; New England Nuclear) added at various time points. Cells were then harvested for RNA isolation and FACS analysis at 0, 1, 2, 4, 8, 16, 24, 28 and 72-hours post androgen addition. FACS analysis was performed using the anti-P501S antibody 10E3- 25 G4-D3 and permeabilized cells.

For Northern analysis, 5-10 micrograms of total RNA was run on a formaldehyde denaturing gel, transferred to Hybond-N nylon membrane (Amersham Pharmacia Biotech, Piscataway, NJ), cross-linked and stained with methylene blue. The filter was then prehybridized with Church's Buffer (250 mM Na_2HPO_4 , 70 mM H_3PO_4 , 30 1 mM EDTA, 1% SDS, 1% BSA in pH 7.2) at 65 °C for 1 hour. P501S DNA was

labeled with 32P using High Prime random-primed DNA labeling kit (Boehringer Mannheim). Unincorporated label was removed using MicroSpin S300-HR columns (Amersham Pharmacia Biotech). The RNA filter was then hybridized with fresh Church's Buffer containing labeled cDNA overnight, washed with 1X SCP (0.1 M NaCl, 0.03 M Na₂HPO₄·7H₂O, 0.001 M Na₂EDTA), 1% sarkosyl (n-lauroylsarcosine) and exposed to X-ray film.

Using both FACS and Northern analysis, P5018 message and protein levels were found to increase in response to androgen treatment.

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EXAMPLE 21

PREPARATION OF FUSION PROTEINS OF PROSTATE-SPECIFIC ANTIGENS

The example describes the preparation of a fusion protein of the prostate-specific antigen P703P and a truncated form of the known prostate antigen PSA. The truncated form of PSA has a 21 amino acid deletion around the active serine site. The expression construct for the fusion protein also has a restriction site at 3' end, immediately prior to the termination codon, to aid in adding cDNA for additional antigens.

The full-length cDNA for PSA was obtained by RT-PCR from a pool of RNA from human prostate tumor tissues using the primers of SEQ ID NO: 607 and 608, and cloned in the vector pCR-Blunt II-TOPO. The resulting cDNA was employed as a template to make two different fragments of PSA by PCR with two sets of primers (SEQ ID NO: 609 and 610; and SEQ ID NO: 611 and 612). The PCR products having the expected size were used as templates to make truncated forms of PSA by PCR with the primers of SEQ ID NO: 611 and 613, which generated PSA (delta 208-218 in amino acids). The cDNA for the mature form of P703P with a 6X histidine tag at the 5' end, was prepared by PCR with P703P and the primers of SEQ ID NO: 614 and 615. The cDNA for the fusion of P703P with the truncated form of PSA (referred to as FOPP) was then obtained by PCR using the modified P703P cDNA and the truncated form of PSA cDNA as templates and the primers of SEQ ID NO: 614 and 615. The FOPP

cDNA was cloned into the NdeI site and XhoI site of the expression vector pCRX1, and confirmed by DNA sequencing. The determined cDNA sequence for the fusion construct FOPP is provided in SEQ ID NO: 616, with the amino acid sequence being provided in SEQ ID NO: 617.

- 5 The fusion FOPP was expressed as a single recombinant protein in *E. coli* as follows. The expression plasmid pCRX1FOPP was transformed into the *E. coli* strain BL21-CodonPlus RIL. The transformant was shown to express FOPP protein upon induction with 1 mM IPTG. The culture of the corresponding expression clone was inoculated into 25 ml LB broth containing 50 ug/ml kanamycin and 34 ug/ml chloramphenicol, grown at 37 °C to OD600 of about 1, and stored at 4 °C overnight.
- 10 The culture was diluted into 1 liter of TB LB containing 50 ug/ml kanamycin and 34 ug/ml chloramphenicol, and grown at 37 °C to OD600 of 0.4. IPTG was added to a final concentration of 1 mM, and the culture was incubated at 30 °C for 3 hours. The cells were pelleted by centrifugation at 5,000 RPM for 8 min. To purify the protein, the
- 15 cell pellet was suspended in 25 ml of 10 mM Tris-Cl pH 8.0, 2mM PMSF, complete protease inhibitor and 15 ug lysozyme. The cells were lysed at 4 °C for 30 minutes, sonicated several times and the lysate centrifuged for 30 minutes at 10,000 x g. The precipitate, which contained the inclusion body, was washed twice with 10 mM Tris-Cl pH 8.0 and 1% CHAPS. The inclusion body was dissolved in 40 ml of 10 mM Tris-Cl
- 20 pH 8.0, 100 mM sodium phosphate and 8 M urea. The solution was bound to 8 ml Ni-NTA (Qiagen) for one hour at room temperature. The mixture was poured into a 25 ml column and washed with 50 ml of 10 mM Tris-Cl pH 6.3, 100 mM sodium phosphate, 0.5% DOC and 8M urea. The bound protein was eluted with 350 mM imidazole, 10 mM Tris-Cl pH 8.0, 100 mM sodium phosphate and 8 M urea. The fractions containing
- 25 FOPP proteins were combined and dialyzed extensively against 10 mM Tris-Cl pH 4.6, aliquoted and stored at -70 °C.

EXAMPLE 22

REAL-TIME PCR CHARACTERIZATION OF THE PROSTATE-SPECIFIC ANTIGEN P501S IN
PERIPHERAL BLOOD OF PROSTATE CANCER PATIENTS

5 Circulating epithelial cells were isolated from fresh blood of normal individuals and metastatic prostate cancer patients, mRNA isolated and cDNA prepared using real-time PCR procedures. Real-time PCR was performed with the TaqmanTM procedure using both gene specific primers and probes to determine the levels of gene expression.

10 Epithelial cells were enriched from blood samples using an immunomagnetic bead separation method (DynaL A.S., Oslo, Norway). Isolated cells were lysed and the magnetic beads removed. The lysate was then processed for poly A+ mRNA isolation using magnetic beads coated with Oligo(dT)₂₅. After washing the beads in buffer, bead/poly A+ RNA samples were suspended in 10 mM Tris HCl pH 8.0
15 and subjected to reversed transcription. The resulting cDNA was subjected to real-time PCR using gene specific primers. Beta-actin content was also determined and used for normalization. Samples with P501S copies greater than the mean of the normal samples + 3 standard deviations were considered positive. Real time PCR on blood samples was performed using the TaqmanTM procedure but extending to 50 cycles using
20 forward and reverse primers and probes specific for P501S. Of the eight samples tested, 6 were positive for P501S and β -actin signal. The remaining 2 samples had no detectable β -actin or P501S. No P501S signal was observed in the four normal blood samples tested.

EXAMPLE 23

EXPRESSION OF THE PROSTATE-SPECIFIC ANTIGENS P703P AND P501S IN
SCID MOUSE-PASSAGED PROSTATE TUMORS

25 When considering the effectiveness of antigens in the treatment of
30 prostate cancer, the continued presence of the antigens in tumors during androgen

ablation therapy is important. The presence of the prostate-specific antigens P703P and P501S in prostate tumor samples grown in SCID mice in the presence of testosterone was evaluated as follows.

Two prostate tumors that had metastasized to the bone were removed from patients, implanted into SCID mice and grown in the presence of testosterone. Tumors were evaluated for mRNA expression of P703P, P501S and PSA using quantitative real time PCR with the SYBR green assay method. Expression of P703P and P501S in a prostate tumor was used as a positive control and the absence in normal intestine and normal heart as negative controls. In both cases, the specific mRNA was present in late passage tumors. Since the bone metastases were grown in the presence of testosterone, this implies that the presence of these genes would not be lost during androgen ablation therapy.

EXAMPLE 24

ANTI-P503S MONOCLONAL ANTIBODY INHIBITS TUMOR GROWTH *IN VIVO*

The ability of the anti-P503S monoclonal antibody 20D4 to suppress tumor formation in mice was examined as follows.

Ten SCID mice were injected subcutaneously with HEK293 cells that expressed P503S. Five mice received 150 micrograms of 20D4 intravenously at day 0 (time of tumor cell injection), day 5 and day 9. Tumor size was measured for 50 days. Of the five animals that received no 20D4, three formed detectable tumors after about 2 weeks which continued to enlarge throughout the study. In contrast, none of the five mice that received 20D4 formed tumors. These results demonstrate that the anti-P503S Mab 20D4 displays potent anti-tumor activity *in vivo*.

EXAMPLE 25

CHARACTERIZATION OF A T CELL RECEPTOR CLONE FROM A P501S-SPECIFIC T CELL CLONE

T cells have a limited lifespan. However, cloning of T cell receptor (TCR) chains and subsequent transfer essentially enables infinite propagation of the T

cell specificity. Cloning of tumor-antigen TCR chains allows the transfer of the specificity into T cells isolated from patients that share the TCR MHC-restricting allele. Such T cells could then be expanded and used in adoptive transfer settings to introduce the tumor antigen specificity into patients carrying tumors that express the antigen. T cell receptor alpha and beta chains from a CD8 T cell clone specific for the prostate-specific antigen P501S were isolated and sequenced as follows.

Total mRNA from 2×10^6 cells from CTL clone 4E5 (described above in Example 12) was isolated using Trizol reagent and cDNA was synthesized. To determine Va and Vb sequences in this clone, a panel of Va and Vb subtype-specific primers was synthesized and used in RT-PCR reactions with cDNA generated from each of the clones. The RT-PCR reactions demonstrated that each of the clones expressed a common Vb sequence that corresponded to the Vb7 subfamily. Furthermore, using cDNA generated from the clone, the Va sequence expressed was determined to be Va6. To clone the full TCR alpha and beta chains from clone 4E5, primers were designed that spanned the initiator and terminator-coding TCR nucleotides. The primers were as follows: TCR Valpha-6 5'(sense): GGATCC---GCCGCCACC---ATGTCACCTTCTAGCCTGCT (SEQ ID NO: 899) BamHI site Kozak TCR alpha sequence TCR alpha 3' (antisense): GTCOAC---TCAGCTGGACCACAGCCGCAG (SEQ ID NO: 900) SalI site TCR alpha constant sequence TCR Vbeta-7. 5'(sense): GGATCC---GCCGCCACC---ATGGGCTGCAGGCTGCTCT (SEQ ID NO: 901) BamHI site Kozak TCR alpha sequence TCR beta 3' (antisense): GTCGAC---TCAGAAATCCCTTCTCTTGAC (SEQ ID NO: 902) SalI site TCR beta constant sequence. Standard 35 cycle RT-PCR reactions were established using cDNA synthesized from the CTL clone and the above primers, employing the proofreading thermostable polymerase PWO (Roche, Nutley, NJ).

The resultant specific bands (approx. 850 bp for alpha and approx. 950 for beta) were ligated into the PCR blunt vector (Invitrogen) and transformed into *E. coli*. *E. coli* transformed with plasmids containing full-length alpha and beta chains were identified, and large scale preparations of the corresponding plasmids were generated. Plasmids containing full-length TCR alpha and beta chains were submitted

for sequencing. The sequencing reactions demonstrated the cloning of full-length TCR alpha and beta chains with the determined cDNA sequences for the Vb and Va chains being shown in SEQ ID NO: 903 and 904, respectively. The corresponding amino acid sequences are shown in SEQ ID NO: 905 and 906, respectively. The Va sequence was shown by nucleotide sequence alignment to be 99% identical (347/348) to Va6.2, and the Vb to be 99% identical to Vb7 (336/338).

EXAMPLE 26

CAPTURE OF PROSTATE SPECIFIC CELLS USING
THE PROSTATE ANTIGEN P503S

As described above, P503S is found on the surface of prostate cells. Secondary coated microsphere beads specific for mouse IgG were coupled with the purified P503S-specific monoclonal antibody 1D12. The bound P503S antibody was then used to capture HEK cells expressing recombinant P503S. This provides a model system for prostate-specific cell capture which may be usefully employed in the detection of prostate cells in blood, and therefore in the detection of prostate cancer.

P503S-transfected HEK cells were harvested and redissolved in wash buffer (PBS, 0.1% BSA, 0.6% sodium citrate) at an appropriate volume to give at least 5^4 cells per sample. Round bottom Eppendorf tubes were used for all procedures involving beads. The stock concentrations were as shown below in Table VIII.

Table VIII

Stock concentrations	Sample concentration	Amount needed
Epithelial enrich beads 4^8 beads/ml (Dyna1 Biotech Inc. Lake Success, NY)	1^7 beads/ml	125 ul stock per 5 ml volume
1D12 ascites antibody 2 mg/ml	0.1 ug/ml (0.1X) to 5 ug/ml (5X) titrations	0.05 ul to 2.5 ul stock per sample
α -Mamma Mu 0.9 mg/ml	1 ug/ml (1X)	1.1 ul stock per sample
Pan-mouse IgG beads 4^8 beads/ml (Dyna1 Biotech)	1^7 beads/ml	125 ul stock per 5 ml volume

Blocked immunomagnetic beads were pre-washed as follows: all beads needed were pooled and washed once with 1 ml wash buffer. The beads were resuspended in a 3X volume of 1% BSA (v/v) in wash buffer and incubated for 15 min rotating at 4 °C. The beads were then washed three times with 2X volume of wash buffer and resuspended to original volume. Non-blocked beads were pooled, washed three times with 2X volume of wash buffer and resuspended to original volume.

Primary antibody was incubated with secondary beads in a fresh Eppendorf for 30 minutes, rotating at 4 °C. Approximately 200 ul wash buffer was added to increase the total volume for even mixing of the sample. The antibody-bead solution was transferred to a fresh Eppendorf, washed twice with an equal volume of wash buffer and resuspended to original volume. Target cells were added to each sample and incubated for 45 minutes, rotating at 4 °C. The tubes were transferred to a magnet, the supernatant removed, taking care not to agitate the beads, and the samples were washed twice with 1 ml wash buffer. The samples were then ready for RT-PCR using a Dynabeads mRNA direct microkit (DynaL Biotech).

Epithelial cell enrichment was placed in a magnet and supernatant was removed. The epithelial enrichment beads were then resuspended in 100 ul lysis/binding buffer fortified with Rnasin (2 U/ul per sample), and stored at -70 °C until use. Oligo (dT₂₅) Dynabeads were pre-washed as follows: all beads needed were pooled (23 ul/sample), washed three times with an excess volume of lysis/binding buffer, and resuspended to original volume. The lysis supernatant was separated with a magnet and transferred to a fresh Eppendorf. 20 ul oligo(dT₂₅) Dynabeads were added per sample and rolled for 5 min at room temperature. Supernatant was separated using a magnet and discarded, leaving the mRNA annealed to the beads. The bead/mRNA complex was washed with buffer and resuspended in cold Tris-HCl.

For RT-PCR, the Tris-HCl supernatant was separated and discarded using MPS. For each sample containing 1⁵ cells or less, the following was added to give a total volume of 30 ul: 14.25 ul H₂O; 1.5 ul BSA; 6 ul first strand buffer; 0.75 mL 10 mM dNTP mix; 3 ul Rnasin; 3 ul 0.1M dTT; and 1.5 ul Superscript II. The resulting solution was incubated for 1 hour at 42 °C, diluted 1:5 in H₂O, heated at 80°C for 2 min

to detach cDNA from the beads, and immediately placed on MPS. The supernatant containing cDNA was transferred to a new tube and stored at -20 °C.

Table IX shows the percentage of capture of P503S-transfected HEK cells as determined by RT-PCR.

5

Table IX

	% capture P503S-transfected HEK cells	% capture LnCAP cells
0.1 ug/ml P503S Mab	36.90	0.00
0.5 ug/ml P503S Mab	67.40	2.93
1 ug/ml P503S Mab	40.22	0.00
5 ug/ml P503S Mab	13.11	0.00
Anti-Mu beads only, non-blocked	1.42	0.00
Anti-Mu beads only, blocked	15.65	20.21
Absolute control, non-capture cells	100.00	100.00

10

From the foregoing it will be appreciated that, although specific embodiments of the invention have been described herein for purposes of illustration, various modifications may be made without deviating from the spirit and scope of the invention. Accordingly, the invention is not limited except as by the appended claims.

CLAIMS

What is Claimed:

1. An isolated polynucleotide comprising a sequence selected from the group consisting of:

(a) sequences provided in SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942;

(b) complements of the sequences provided in SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942;

(c) sequences consisting of at least 20 contiguous residues of a sequence provided in SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942;

(d) sequences that hybridize to a sequence provided in SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942 under moderately stringent conditions;

(e) sequences having at least 75% identity to a sequence of SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942;

(f) sequences having at least 90% identity to a sequence of SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942; and

(g) degenerate variants of a sequence provided in SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591, 593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942.

2. An isolated polypeptide comprising an amino acid sequence selected from the group consisting of:

(a) sequences recited in SEQ ID NO: 112-114, 172, 176, 178, 327, 329, 331, 336, 339, 376-380, 383, 477-483, 496, 504, 505, 519, 520, 522, 525, 527, 532, 534, 537-551, 553-568, 573-586, 588-590, 592, 706-708, 775, 776, 778, 780, 781, 811, 814, 818, 826, 827, 853, 855, 858, 860-862, 866-877, 879, 883-893, 895, 897, 898, 909-915, 920-928, 932-934, 940, 941 and 943;

(b) sequences having at least 70% identity to a sequence of SEQ ID NO: 112-114, 172, 176, 178, 327, 329, 331, 336, 339, 376-380, 383, 477-483, 496, 504, 505, 519, 520, 522, 525, 527, 532, 534, 537-551, 553-568, 573-586, 588-590, 592, 706-708, 775, 776, 778, 780, 781, 811, 814, 818, 826, 827, 853, 855, 858, 860-862, 866-877, 879, 883-893, 895, 897, 898, 909-915, 920-928, 932-934, 940, 941 and 943;

(c) sequences having at least 90% identity to a sequence of SEQ ID NO: 112-114, 172, 176, 178, 327, 329, 331, 336, 339, 376-380, 383, 477-483, 496, 504, 505, 519, 520, 522, 525, 527, 532, 534, 537-551, 553-568, 573-586, 588-590, 592, 706-708, 775, 776, 778, 780, 781, 811, 814, 818, 826, 827, 853, 855, 858, 860-862, 866-877, 879, 883-893, 895, 897, 898, 909-915, 920-928, 932-934, 940, 941 and 943;

(d) sequences encoded by a polynucleotide of claim 1;

(e) sequences having at least 70% identity to a sequence encoded by a polynucleotide of claim 1; and

(f) sequences having at least 90% identity to a sequence encoded by a polynucleotide of claim 1.

3. An expression vector comprising a polynucleotide of claim 1 operably linked to an expression control sequence.

4. A host cell transformed or transfected with an expression vector according to claim 3.

5. An isolated antibody, or antigen-binding fragment thereof, that specifically binds to a polypeptide of claim 2.

6. A method for detecting the presence of a cancer in a patient, comprising the steps of:

(a) obtaining a biological sample from the patient;

(b) contacting the biological sample with a binding agent that binds to a polypeptide of claim 2;

(c) detecting in the sample an amount of polypeptide that binds to the binding agent; and

(d) comparing the amount of polypeptide to a predetermined cut-off value and therefrom determining the presence of a cancer in the patient.

7. A fusion protein comprising at least one polypeptide according to claim 2.

8. An oligonucleotide that hybridizes to a sequence recited in SEQ ID NO: 1-111, 115-171, 173-175, 177, 179-305, 307-315, 326, 328, 330, 332-335, 340-375, 381, 382 and 384-476, 524, 526, 530, 531, 533, 535, 536, 552, 569-572, 587, 591,

593-606, 618-705, 709-774, 777, 789, 817, 823, 824, 878, 880-882, 894, 896, 907, 908, 916-919, 929-931, 938, 939 and 942 under moderately stringent conditions.

9. A method for stimulating and/or expanding T cells specific for a tumor protein, comprising contacting T cells with at least one component selected from the group consisting of:

- (a) polypeptides according to claim 2;
- (b) polynucleotides according to claim 1; and
- (c) antigen-presenting cells that express a polypeptide according to claim 2,

under conditions and for a time sufficient to permit the stimulation and/or expansion of T cells.

10. An isolated T cell population, comprising T cells prepared according to the method of claim 9.

11. A composition comprising a first component selected from the group consisting of physiologically acceptable carriers and immunostimulants, and a second component selected from the group consisting of:

- (a) polypeptides according to claim 2;
- (b) polynucleotides according to claim 1;
- (c) antibodies according to claim 5;
- (d) fusion proteins according to claim 7;
- (e) T cell populations according to claim 10; and
- (f) antigen presenting cells that express a polypeptide according to claim 2.

12. A method for stimulating an immune response in a patient, comprising administering to the patient a composition of claim 11.

13. A method for the treatment of a cancer in a patient, comprising administering to the patient a composition of claim 11.

14. A method for determining the presence of a cancer in a patient, comprising the steps of:

- (a) obtaining a biological sample from the patient;
- (b) contacting the biological sample with an oligonucleotide according to claim 8;

- (c) detecting in the sample an amount of a polynucleotide that hybridizes to the oligonucleotide; and

- (d) compare the amount of polynucleotide that hybridizes to the oligonucleotide to a predetermined cut-off value, and therefrom determining the presence of the cancer in the patient.

15. A diagnostic kit comprising at least one oligonucleotide according to claim 8.

16. A diagnostic kit comprising at least one antibody according to claim 5 and a detection reagent, wherein the detection reagent comprises a reporter group.

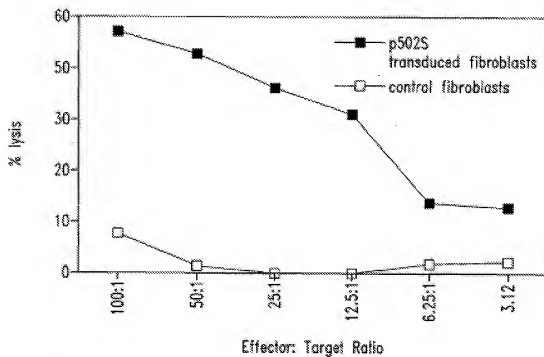
17. A method for inhibiting the development of a cancer in a patient, comprising the steps of:

- (a) incubating CD4+ and/or CD8+ T cells isolated from a patient with at least one component selected from the group consisting of: (i) polypeptides according to claim 2; (ii) polynucleotides according to claim 1; and (iii) antigen presenting cells that express a polypeptide of claim 2, such that T cell proliferate;

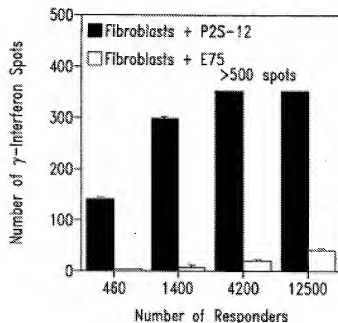
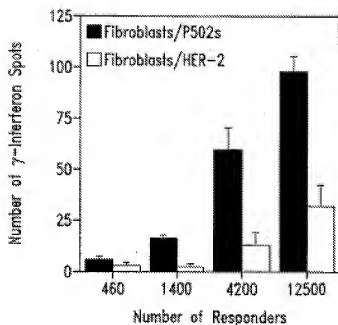
- (b) administering to the patient an effective amount of the proliferated T cells,

and thereby inhibiting the development of a cancer in the patient.

1/14

*Fig. 1*

2/14

*Fig. 2A**Fig. 2B*

3/14

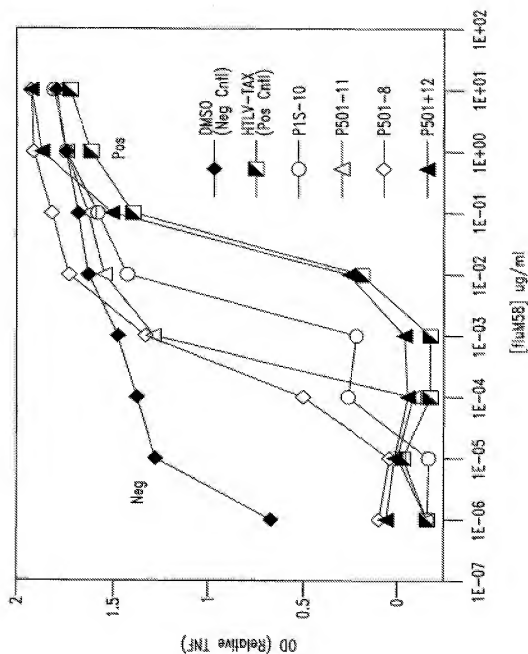
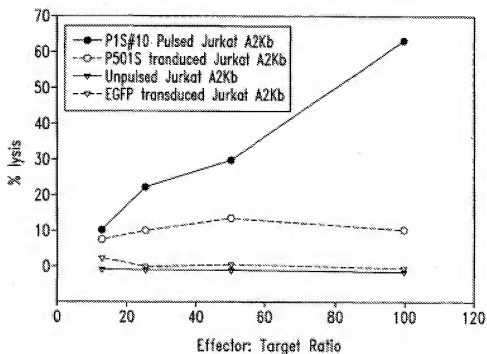
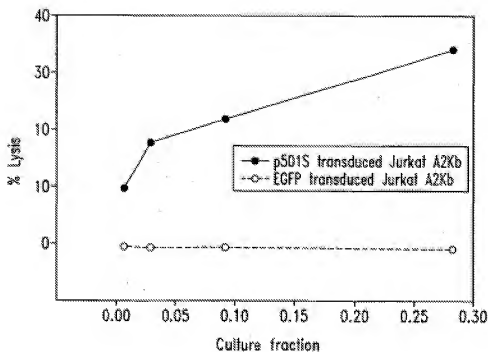
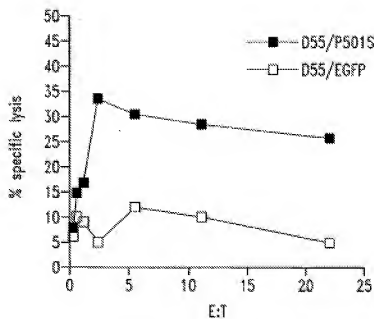
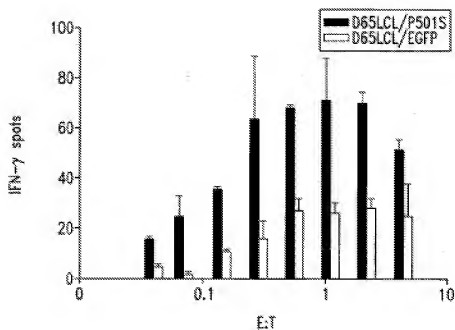


Fig. 3

4/14

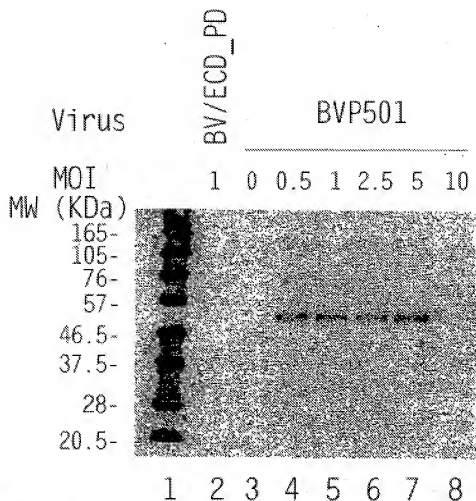
*Fig. 4**Fig. 5*

5/14

*Fig. 6A**Fig. 6B*

6/14

Expression of P501S
by the Baculovirus Expression System



0.6 million high 5 cells in 6-well plate were infected with an unrelated control virus BV/ECD_PD (lane2), without virus (lane3), or with recombinant baculovirus for P501 at different MOIs (lane 4-8). Cell lysates were run on SDS-PAGE under the reducing conditions and analyzed by Western blot with a monoclonal antibody against P501S (P501S-10E3-G4D3). Lane 1 is the biotinylated protein molecular weight marker (BioLabs).

Fig. 7.

7/14

FIGURE 8. Mapping of the epitope recognized by 10E3-G4-D3

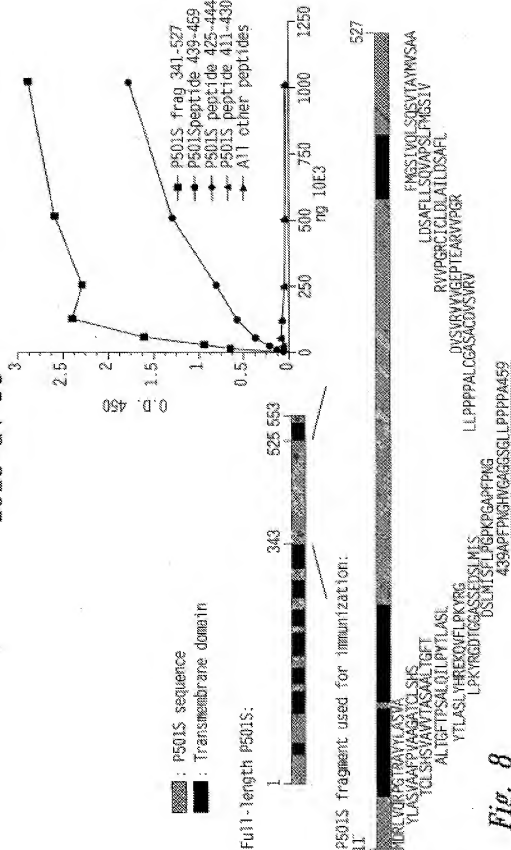


Fig. 8

8/14

Schematic of P501S with predicted
transmembrane, cytoplasmic, and extracellular regions

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DHWGRGYGRRRP FIWALSLGILLSLFLIPRAGWL AGLLCPDPRPLE LALLILGVGLLDFCQGVCFTPL

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HQLCCRMPTLRR LFVAELCSWMALMTFTLFYTDF VGEGLYQGVPRAEPGTEARRHYDEGVR

MGSLGLFLQCAISLVFSLVM DRLVQRFGTRAVYLAS VAAFPVAAGATCLSHSVAVVTA SAA

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LPPPPALCGASACDVSVRVVVGEPTEARVVPGRG ICLDLAILDSAFLLSQVAPSLF MGSIVQLSQS

VTAYMVSAAGLGLVATYFAT QVVFDKSDLAKYSA

Underlined sequence: Predicted transmembrane domain; Bold sequence:
Predicted extracellular domain; *Italic sequence*: Predicted intracellular
domain. Sequence in bold/underlined: used generate polyclonal rabbit
serum

Localization of domains predicted using HMMTOP (G.E. Tusnady and I. Simon
(1998) Principles Governing Amino Acid Composition of Integral Membrane
Proteins: Applications to topology Prediction. J.Mol Biol. 283, 489-506.

Fig. 9

9/14

Genomic Map of (5) Corixa Candidate Genes

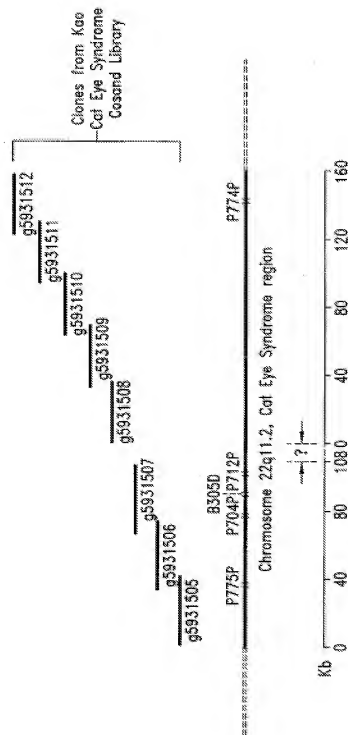


Fig. 10

10/14

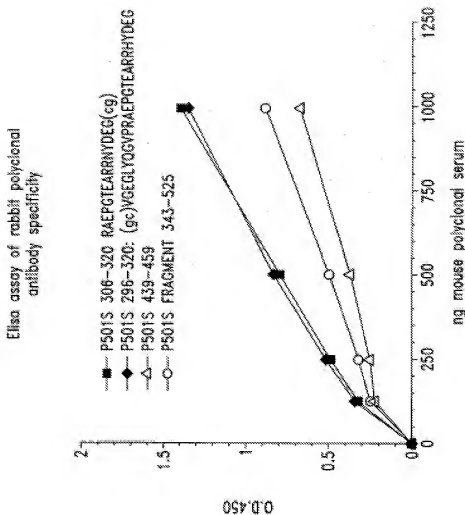


Fig. 11

11/14

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Fig. 12A (1)

12/14

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Fig. 12A (2)

13/14

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Fig. 12A (3)

14/14

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Fig. 12B

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 Mitcham, Jennifer L.
 Harlocker, Susan L.
 Yuqiu, Jiang
 Kalos, Michael D.
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 Stolk, John A.
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 tccctgtcca tctgattgca aagttcatca gacttttagc cahmctcttt gatcaagcagc 300
 tctgagaact ggggttctat tgcctcaaca gccatgaatt ccccatctcg tgtcctgtaa 360
 gtgctataga aagggtgctcc accatccaac atgttctgtc ctgcaggggg ggcocgggtac 420
 ccaattctgc ctatantgag tctgtattac cgcgctcact ggcgctctgt ttacaacgtc 480
 gtgactggga aaacccctgg cgttaaccaac ttaactgcct tgcagacact cccocctttcg 540
 ccagctgggg gtaatanaga aaagggccgc accgatggcc ctccaacagc ttgcgcacct 600
 gaatgggnaa atgggacccc cctgtttacc cgcatttaac ccccgcnagg ttctngttgtt 660
 accccacact nnaacgctta caatttgcaa gcgccttanc gwcgctccc ttctnctctt 720
 ctctcccttcc ttctnctnccn ctttcccccg gggtttcccc cntcaaaacc ona 773

<210> 4
 <211> 828
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(828)
 <223> n = A,T,C or G

<400> 4

oactctgagt	oactatgacc	tgtgctttct	ggtgtggagt	coagggctgc	taggaaaaagg	60
aatgggcaga	cacaggtgta	tgccaatggt	tctgaatggt	gtataatttc	gtcctctcct	120
toggiaacat	ggctgtctct	gaagacttnt	cgctcagttt	cagtgaggac	acacacaaag	180
acgttgggtga	ccatgttgtt	tgtgggtgct	agagatggga	gggttggggc	ccacccctgga	240
agagtggaca	gtgacacaa	gtggacactc	tctacagatc	actgaggata	agctggagcc	300
acastgcagt	agggcacacac	acagcaagga	tgaacctgta	aacatagccc	acgtgtctct	360
gnggggcactg	gggaagctat	atnaggccgt	gagcanaaag	asggggagga	tccactagtt	420
ctanaagcggc	cgccacggcg	gtgganctnc	ancttttgtt	ccctttagtg	agggtttaatt	480
ggcgctctgg	ontaactctg	gtctatnctn	tttctgtgtg	gaattgttta	tccgtctaca	540
attccacaca	acatacganc	gggaaccata	aantgtaaac	ctgggtgtcc	taatgamtga	600
ctaatctaca	ttaattgggt	tgogctcaat	gcccgttttc	caatonggaa	acctgtcttg	660
ccacttgcat	taattgcatn	gcacaccccc	ggggaanaag	gtttgctgtt	tgggcgctct	720
tcogcttctc	cnctcanta	ntccctnccc	tgggtctctc	cggtgcgncg	aaacccgttc	780
acnccctoca	aagggggtat	tccgttttcc	ccnaatccgg	ggananc		828

<210> 5
 <211> 834
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(834)
 <223> n = A,T,C or G

<400> 5

tttttttttt	tttttactga	tagatggaat	ttattaagct	tttccactgt	gatagccat	60
agtfttaatt	gcattccaaag	tactatacaaa	aactctagca	atcaagaatg	gcagcatggt	120
attttataac	aatacacaac	tgtggctttt	aaaatttgggt	tttctaaaga	taatttatac	180
tgaagttaac	ctagcccatgc	ttttaaaaaa	tgttttaggt	cactccaagc	ttggcagttt	240
acatttggca	taaacataaa	taaaacacac	acaatttaac	aaatacaaaa	tacaacattg	300
taggcataaa	tactatacag	tataaggaaa	aggtggttagt	gttgagttaag	cagttatfatg	360
aatagaattac	cttggcctct	atgcaaatat	gtctagacac	tttgattcac	tcagccctga	420
cattcagttt	tccaagttagg	agacaggttc	tacagtatca	ttttacagtt	tccaacacat	480
tgaanaacaa	tgaasaatga	tgaatttgatt	ttttattaatg	cattacatcc	tcaagagttta	540
tcaccacccc	ctcagtttata	aaaaattttc	asgttatatt	agtcataaaa	cttgggtgtgc	600
ttattttaaa	ttagtgtctaa	atgatttaag	tgaagacaa	aattgtcccc	taattgtgatt	660
gatatttgto	atttttaacc	gottctaaat	ctnaactttc	aggtttttga	tggtgaacat	720
tgnatnana	tgttccanag	tincaaccta	ctggaacatt	acagttgtgt	actgtcaaaa	780
tgttattttg	tttaaatatta	aattttaacc	tggtggaaaa	ataattttaa	atna	834

<210> 6
 <211> 818
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(818)
 <223> n = A,T,C or G

<400> 6

```

tttttttttt tttttttttt aagaccctca tcaatagatg gagacataca gaastagtcn 60
aacccatctc acaaaatgco agtatcagcg ggcggcttcg aagccaaagt gatgtttgga 120
tgtaaagtga satattagtt ggcggatgaa gcagatagtg aggaagttg agccaataat 180
gagctgaagt cagtggaaag ctgtggctac aaaaaatgtt gagccgtaga tcccgtcgga 240
aatgttgaag ggagactcga agtactctga ggcctgttag agggtaaaat agagacccag 300
taaaatttga ataaagcagt ctgnaattat ttggtttcgg ttgttttcta ttgactatcg 360
gtgacttcga gtgattgata ctccgatgc gagttaacg gatgtgttta ggagtgggac 420
ttctagggga tttagcgggg tgatgcctct tggggggcag tgcctccta gttggggggg 480
aggggctagg ctggagtggt aaaaagctca gaaaaatcct gcgaagaaaa aaacttctga 540
ggtaataaat aggaattatcc cgtatcgaag gccttttttg acaggttgghg tgtggtggcc 600
ttggtatgtg ttctctcgtg ttacatcgcg ccatcattgg tatagtgta gtgtgtttgg 660
ttantanggc ctantatgaa gaacttttgg antggaatta aatcaatnge ttggccggaa 720
gtcattanga ngcctnaaaa ggcctgtta aggtctctgg ctnggttktta cccaaoccat 780
ggaatacaca ccccggaana ntgnatccct attcttaa 818

```

<210> 7

<211> 817

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(817)

<223> n = A, T, C or G

<400> 7

```

tttttttttt tttttttttt tggctctaga gggggtagag ggggtgtgat agggtaata 60
cgggcoctat ttcasagatt tttagggaa ttaattctag gcuytggggt atgaascctg 120
ggttgtctoc acagailkca gagcattgac cgtagtatac ccocpgtgcg gtacgggtga 180
aagtgttttg gtttagacyt ccgggaattg catctgtttt kaagcotaat gtggggacag 240
ctcatgagtg cagagagctct tgtgatgtaa ttattataca satgggggct tcaatcggga 300
gtactactcyg attgtcaacg tcaaggagtc gcaggtcgcc tggttctagg aataatgggg 360
gaagtatgta ggaattgagag attaatccgc cgtagtccgt gttctcctag gtccaatacc 420
attgtgtccc aattgtattg atggtaaggg gagggatcgt tgaactcgto tgttatgtaa 480
aggatnccct ngggatggga aggcnaftnaa ggaactangga tnaatggcgg gcangataat 540
tcaaaongtc tctantctct gaacgctctg aaatgttaat aanaattaan ttngttatt 600
gaatttngn gaaaagggct taccggacta gaaaccaaat angaasanta atnntasag 660
cmttatonta aaaggtlnata accnctccta tnatcccaac catngnatt cccacnccnn 720
acnattggat nccocanttc canasaggcg cncoccccg tgnannccnc ctittgttcc 780
cttnantgan ggttattenc coctngentt atcaance 817

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<210> 8

<211> 799

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(799)

<223> n = A, T, C or G

<400> 8

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catttccggg ttactttctt aaggaaagcc gagcgggaag tgcataactg ggaatcggtg 60
cataagagga actttctgct ggcacgcgct agggcaagc ggagagagca ctocagagct 120
ctgaagcgca cgtcccagaa ggtggacttg gcactgaac agctgggaca catcccgag 180
tacgaacagc gctgaagagt gctggagcgg gaggtccagc agtgtagccg cgtcctgggg 240
tgggtggccc angcctganc cgtctgctct tgcctccccc angtggcccg ccccccctcg 300
acctgcctgg gtccaaacac tgagccctgc tggnggactt caagganaac ccccccangg 360

```

```

gggttttggct gctanantaa ggcctatctg ggcctgggac ccccccctg gttggccttg 420
tcttttgagt gagccccatg tccatctggg ccaatgtong gaccaccttt ngggagtgtt 480
ctcctataaa ccaacannatg cccggctact cccggaaacc antccancc tngaaaggtat 540
caagncctcg atccactnnt nctanaaacg gccnccnccg cngtggaaac cncctntnct 600
tccttttnt tnaaggtttaa tnnccgcttg gcttaccan ngtcctnccn ntcttcnnt 660
gttnaaattg ttangcnccc nccntcccn cnaacnncan cccgacccna anntnncan 720
nctgggggt nccnncngat tgcacccncc nccntntant tgcnttnggg naacnngccc 780
cttccctct nggganncg

```

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<210> 9
<211> 801
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(801)
<223> n = A,T,C or G

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```

<400> 9
acgccttgat cctccacggc tgggaactgg tctgggagga gccgggcatg ctgtggtttg 60
taangatgac actcccaaaag gtggtctctga cagtggccca gatggacatg gggctccct 120
caaggcaag gccaccaggt gccggggccg aagccacat gatccttact ctatgagcaa 180
aatccctgt gggggcttct ccttgaagtc cgcacacagg gctcagtctt tggacccang 240
caggtctatg ggtttctnnc caactggggg cncacacgca aaaggccnca gggcctcngn 300
caccatccc angacggcgc tacactnctg gactccncc bccaccaact teatggctg 360
tctnatacgg cgnatnctg ccactgttt cngtgcacac tccancttct nggaagtggc 420
ctactatgc cgggattenc nctccggctt tgtccctatc caagtncan caacaaatt 480
cncctantg caccattcc caonittnnc agnttccncc ancgagcttc ctntaaag 540
ggttganccc cggaaaatnc cccaaagggg gggggccngg taccacaatn cccctnata 600
gctgaatcc ccaathacnn gntcnatgg anccntccnt tttaannnc ttctnaactl 660
gggaananc ctcgncctn ccccnctaa tccncccttg cnaangnnc ccccnctcc 720
nccnncatg gcntnanc nnaaaaggc cnaanncaa tctcctnccn cctccttgg 780
ccnccctgg aatggcccn c
801

```

```

<210> 10
<211> 789
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(789)
<223> n = A,T,C or G

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```

<400> 10
cagcttatnt ggcagctgtg gaagctttcc ctgtggctgc cggtgccaca tgcctgtccc 60
acagttggc cgtggtgaca gttccagccg cctccaccg gttcccttc tccagccctgc 120
agatctctcc ctacacactg gactcctctt cccacccggga gaaagcaggtg ttctctcnc 180
aataccaggy ggaactctga ggtctagca gtaggacag cctgatgacc agcttctgc 240
caggccctaa gcttgagctt ccttcccta atggacacgt ggtgtgtgga ggaagtggcc 300
tctcccccac tccacccggc ctctggcggg cctctgcctg tgatgtccc gtaagtgtgg 360
tggtgggtga gccacccgan gccagggttg ttccggcccg gggcatctgc ctggacctgc 420
ccatcctgga tagtgtcttc tctgtccca ngtgccccc tccctgttta tgggctccat 480
tgtccagctc agccagcttg tcaatgccta tatggtgtct gccgacggcc tgggtctggt 540
ccatttact ttgtacaca ggtantattt gacaaagacc anltggcaca atactcagcc 600
ttaaaaaatt ccagcaaat tgggggttga aggcctggct cactgggtcc aactccccc 660
tcctgttaac ccatggggg tgcgggctg gccgcacatt tctgttgtg ccaantaa 720

```

gtggtctctct gctgccacct gttgctggct gaagtgcnta cngcncanct aggggggtng 780
ggpgttccc 789

<210> 11
<211> 772
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1) ... (772)
<223> n = A, T, C or G

<400> 11
cccccctac ccaatatatta gacaccaaca cagaaasagct agcaatggat tccctcttac 60
tttgttaaat aaataagtta aatatttaaa tgcctgtgtc tctgtgatgg caacagaagg 120
acaaacagge cacatcctga taaaaggtaa ggggggggtg gatcagcaaa aagcagctgc 180
tgtgggctga ggggaacctgg ttcttgtgtg ttgccccca ggcctcttcc cctacaataa 240
actttcataat gtccaatccc catggaggag tgtttcatcc tagaaactcc catgcaagag 300
ctacattaaa cgaagctgca ggttaagggg ctlanayag ggaaccagg tgactgagtt 360
tattcagctc ccaaaaaacc ttctctaggt gtgtctcaac taggaggcta gctgttaacc 420
ctggcctctgg gtaalccacc tgcagagtcu ccgcattcca gtgcattgaa cctttctggc 480
ctccctgtat aagttccagac tgaaccctcc ttggaaggnc tccagtcagg cagccttana 540
aactggggaa aaaaagaaaag gacgccccan ccccagctg tgcantcag caccctasca 600
gacaggggtg gcagcaaaaa aaccacttta ctttggcaca aacaaaaact ngggggggda 660
accctggcac ccnagggggg gtttaacagg ancnngggnaa cntgggaacc aattnaggca 720
ggccnccacc cchaaatntt gctgggaaat ttttctccc ctaaattntt tc 772

<210> 12
<211> 751
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1) ... (751)
<223> n = A, T, C or G

<400> 12
gcaccaatcc cagctggcac acccccacg gtgactgcat tagttcggat gtcatacaaa 60
agctgattga agcaaccctc taccttttgg tegtgaacct ttgtcttggg ccaggtttcca 120
ttggctgtgt tggtagcctt gtccattgcaa cagaatgggg gaagggcaact gttctcttgg 180
aagtaagtg agtccctcaaa atccgtatag ttggtgaag caccagcaact gagccctttc 240
atgggtgtgt tccacacttg agtgaagctc tccctgggaac cataatcttt ctgtatggca 300
ggccctacca gcaactcag ggaagtgctc agccattgtg gtgtacacca aggcagaccac 360
agcagctgcn accctcagaa tgaagatgan gaggongatg aagaagaacg tcnogagggc 420
anactgtgct tcagtcttan caccatanaa gcccttgaaa acccaanaca aagaacccna 480
cnctggctgc gctgaagaaa tnaacccnag ttgacaaact tgcatggcac tggganccac 540
agtggcccca aaatcttcca aaaggatgc cccatcnatt gaccccccaa atgcccactg 600
ccaacagggg ctgcccnaen cnennacga tganconatt gnacaagatc tcnctgtgtt 660
thainaact gaacctgcn tngtggctcc tgttcaggnc cngggcctga cttctnaann 720
aangaactcn gaagncacca cngganann g 751

<210> 13
<211> 729
<212> DNA
<213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(729)
 <223> n = A,T,C or G

<400> 13
 gagccaggag tccctctgac tgcccacatca gtggcaaacac caggaggctg tttgtcctt 60
 tgtggancct cagcagthcc ctcttccaga actcactgac aaganccctg aacaggagcc 120
 accatgcagt gcttcagctt caataagacc atgatgatcc tcttcaattt gctcatcttt 180
 ctgtgtgggt cagccctggt ggcagtgggc atctgggtgt caatogatgg ggcactcttt 240
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 ctcatgcagc cggcgctgtt ggtctttagct ctagggttcc tgggtgtgta tgggtctaaq 360
 actgagagca agtctgacct cgtgacgttc ctcttcaacc tctctctcat ctctcttgc 420
 gaggttgcaa tgcctgggac gcttgggtgt acanccanct ggcctgagca tctctgagct 480
 gttggaacc caccatgaaa gggctcaagt gctgtggcct cncaccaata tcaactcaat 540
 gaagantcac ctacttcaaa gaaaanagtg ccttccccc atttctgtg caattgacaa 600
 acgtccccc caagcccaat tgaacacctg caccccaacc aaatgggttc ccaaccanaa 660
 attcaaggg 720
 729

<210> 14
 <211> 816
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(816)
 <223> n = A,T,C or G

<400> 14
 tgcctctctt caaagtgttt ctgttgcca taacaacac cataggtaaa ggggggcag 60
 tgttcyctga aggggttcta gtaccagcgc gggatgctct ccttgcagag tctgtgtct 120
 ggcagggtcca cgcagtccc ttgtctacgt gggaaatgga tgcgtctggag ctctctcaag 180
 ccaactgtgt atttttcaaa ggcagcctcg tccgaagcgt cggggcagtt ggggtgtct 240
 tcaactctca ggaactgtc natgcagcag ccattgtctc agcggaactg ggtgggtctg 300
 caagtgcagc agcacactgg atggcgctt tccatgnnan gggccctgng ggaagtctcc 360
 tganccocan anctgcctct caaangcccc accttgcaca ccccgacagg ctagaatgga 420
 atctctctcc cgaaggttag ttctcttctg tgcccaancc anccocntaa acaaactctt 480
 gcanatctgc tccggggggg tontantacc ancggtggaa agaaacccca ggmccgagac 540
 caancttgtt tggctnccaa gonataatct nctnttctgc ttggtggaa gcaacantna 600
 ctgtnnamct ttagnccmgt gtctctatgg gttgmnottg aactaatacn ccnstcaast 660
 gggcaaggt aantagcmt ccttnaatt cccanantcn cccctgtgtt tgggtgtttn 720
 cncctcta cccagaaan nccgtgttcc ccccaacta ggggcnaaa ccnntnttcc 780
 cacaacctta ccccaaccc ggggtcngt ggttng 816

<210> 15
 <211> 783
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(783)
 <223> n = A,T,C or G

<400> 15
 ccaagccctg ggcaggcata naattgaagg taacaaccca ggaacccctg gtgctgaagg 60

atgttgaaaa	cacagattgg	cgactactgc	gggttgacac	ggatgtcagg	gtagagagga	120
aagacccaaa	ccaggtggaa	ctgtggggac	tcaaggaang	cacctactgt	tccagatga	180
cagtactag	ctcagancac	ccagaggaca	cggccacagt	cacagtcact	gtgctgtcca	240
ccaaagcagc	agaagactac	tgccttgcac	ccaacaangt	ggtgtgctgc	cggggctctt	300
tccaaocgtg	gtactatgac	cccacggagc	agatctgcac	gagtttctgt	tatggagggt	360
gcttggggcaa	caagaacac	taccttgggg	aagaagagtg	cattctanc	gttctgggtg	420
tgaagggtgg	gcctttgana	ngcactctg	gggtctcang	gactttcccc	caggggccct	480
ccatgggaag	ggcccatcca	ntgttctctg	gcacctgtca	gcccaacccg	tcccgctgca	540
ncaatggctg	ctgcactcac	antttctctg	aattgtgaca	acacccccca	ntgcccccaa	600
ccctcccaac	aaagcttccc	tgtttaaana	tacnccantt	ggtttttnac	aaaccccggg	660
cnctcctntt	ttcccctntn	aacaaagggc	ncctngcctt	gaactgccc	aaacccggaa	720
tctnccnngg	aaaaantccc	ccccctgggt	cctnaaanc	cctccncaa	aaactcccc	780
ccc						783

<210> 16

<211> 801

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(801)

<223> n = A, T, C or G

<400> 16

gcccacattc	cagctgmac	accacccacy	gtgactgcat	tagttcggat	gtcatacaaa	60
agctgattga	agcaacccct	tactttttgg	togtgagcct	tttctgttgg	gcagggttca	120
ttggtgtgtg	tgggtgacgt	gtcatttgcaa	cagaatgggg	gaaaggaact	gttctctctt	180
aagtaggggt	agtcctcaaa	atccgtatag	tgggtgaagc	cacagcaact	gagccctttc	240
atggttggtg	tccacacttg	agtgaagtct	tccctgggac	cataatcttt	cttgatggca	300
ggcactasca	gcncgttcag	gaagtgtcca	gccattgtgg	tgtacaocaa	gycgaccaca	360
gcagctgcac	cctcagcaat	gaagtatggg	aggaggttga	agaagaacgt	cncgagggca	420
cacttgcctc	cgtctttagc	accatagcag	cccangaaac	caagagcaaa	gaccaacaag	480
ccngctgcga	atgaagaana	ntaccccact	tgacaaactg	catggccact	ggacgacagt	540
tggcccgaa	atcttcagaa	aagggatgcc	ccatcgattg	aacacccana	tgcccaactg	600
cnacagggct	gcncnccncc	gaaagaatga	gcaattgaag	saggaatctc	ntgttcttaa	660
tgaactgaaa	ccntgcattg	tggccctctg	tcagggtctc	tggcagtgaa	ttctganaaa	720
aaggaaacgc	ntahagcccc	ccaaagana	aaacaccccc	gggtgttgcc	ctgaatttgc	780
ggccaaggan	ccctgcccnn	g				801

<210> 17

<211> 740

<212> DNA

<213> Homo sapien

<220>

<221> misc_features

<222> (1)...(740)

<223> n = A, T, C or G

<400> 17

gtgagagcca	aggctccctc	tgcctgccc	ctcagtgcca	acacccggga	gctgttttgt	60
cttttctgga	gcctcagcag	ttccctcttt	cagaactcac	tgccaaagac	cttgaaacag	120
agccacccatg	cagtgcttca	gcttcattaa	gaacatgatg	atcctcttca	atttgtctat	180
ctttctgtgt	ggtgcagccc	tgttggcagt	gggcatctgg	gtgtcaatcg	atggggccac	240
ctttctgaag	atcttctggc	cactgtctgc	cagtgccatg	cagtttctga	acgtgggcta	300
cttctctatc	gcagccggcg	tgtgtgtctt	tgtctctgtt	tctctgggct	ctctatgtgc	360
taagacggag	agcaagtgtg	ccctgtgac	gttctctctc	atctctctcc	tcattctcat	420

tgctgaagtt	gcagctgctg	tggtgcctt	ggtgtacac	acaaaggctg	aacattctct	480
gaagttgctg	gtantgctg	ccatcaanaa	agattatggg	ttcccaaggaa	aatttcactc	540
aanmtgga	caacnccatg	aaaaaggctc	caattttctg	tggtttccoc	aactatacgg	600
gaattttgaa	agantccccc	taattccaaa	aaaaaanani	tgctttttcc	ccctttctgt	660
tgcaatgaa	acntcccaan	acagcccaatn	aaaactgtcc	cnanccaaaa	ggntcccaaa	720
caaaaaaant	nnaaggggtt					740

<210> 18

<211> 802

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(802)

<223> n = A,T,C or G

<406> 18

ccgctgggttg	cgctggctcc	gngnagccac	gaagcacgtc	agcatcacca	gcotcaalca	60
caaggtcttc	cagctgcgcg	acattacgca	gggcaagagc	ctccagcaac	actgcattatg	120
gggtacacatt	tactttagca	gcccagggtga	caactgagag	gtgtcgagagc	ttattctctct	180
gagctcttgt	tagtgaggga	agattccggg	cttcagctaa	gtagtcagcg	tatgtcccat	240
aagcaaacac	tgtagcagcg	cggaaggtag	sggcaaaatc	actctcagcg	agctctctaa	300
cattggagcat	gtccagcagt	ctccaaaaca	cgtagacacc	agnggcctcc	agcaactgat	360
ggatgaggtg	ggccacgcgt	gcccccttgg	cgaacttgcc	taggagcaga	aattgtctct	420
ggttctggcc	tgtaacttc	acttcgcgac	tcactcaetg	actgagtggtg	ggggacttgg	480
gtccagagtg	tcagagagcg	tggttcggcc	ccctcactta	atgacacccg	ccanncaacc	540
gtcgctccoc	gcgcantgny	ttcgtctgtc	ctgggtccgg	gtctgtgtggc	cnctactctg	600
aanctctgtc	nggcacatgy	aattccccc	acgggaactn	gtanqatcca	ctmtttctat	660
aacccgagcg	cacccgcnml	gsaactccac	tcitnttccc	tttacttgag	ggttaagctg	720
accctttncg	ttaccttggg	ccaaacccnt	ccntgtgtcg	analgntnaa	tcnggnccna	780
ttcccacccc	atangaagcc	ng				802

<210> 18

<211> 731

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(731)

<223> n = A,T,C or G

<406> 19

cnaagcttcc	aggttaacggg	cgcnaaaccc	tgaccnagg	tancanaaag	cagncgcggg	60
gagccacacg	tcacagagng	ngtcttttat	nggagggggg	ggagccacat	cnctggccat	120
cnlgecccca	actcccccnc	ncncantgca	gtgatgagtg	cagaaactga	ggtnacgttg	180
caggacccaa	gancaaannc	tgctccmgtc	caagtgccga	nagggggggg	ggctggccac	240
cgccctccnt	cnagtgtctg	aaagcccccnn	ccgtgtact	tggtttggaga	acngcnnaag	300
catgccacgn	gttanataac	ngcngagag	taantttgoc	tctcccttcc	ggctggccan	360
cgnrnttctg	tagnggacat	aacttgacta	cttaactgaa	cccnngaatc	tnccnccctt	420
ccactaaagt	cagaaacaana	aacttogaca	ccactcaant	gtccactgnc	tgctcaagta	480
aagtctaccc	catncccaat	gtntgctnga	ngctctgnoc	tgcnttngt	tcggctctgg	540
gaagacccat	caattnaagc	tatgtttctg	actgcctctt	gctccctgna	acaanccncc	600
cnncnntcca	aggggggggnc	ggcccccacat	cccccccaac	ntnaatttna	tttanccccc	660
ccccnngccc	ggccttttta	cnancntann	nnacnnggna	saacennnag	tttccccca	720
nnaatccncc	t					731


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<210> 20
<211> 754
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)... (754)
<223> n = A, T, C or G

<400> 20
tttttttttt tttttttttt taaaaacccc ctccatttna tgnaaacttc cgaattgttc      60
caaccaccctc nccccaaatn ccttttcggg gnggggggttc caaacccaan ttanntttgg      120
aannttaaat aaatntntnt tggngynnaa anccnaatgt nangaaagt naaccacanta      180
tnanetttna tncctggaaa cngtngntt ccaaaaatnt ttaaccctta antccctcgg      240
aaatngttta nggaaaaccc aantctcctt aaggttggtt gaaggatnaa tnaaaanccc      300
smccaatgtt ttttngccac gcttgaatta atttgnttcc gntgttttcc attaaaaana      360
gynnancccc ggttantaan tccccccnnc cccaattata ccganttttt ttngaattgg      420
gancocccgg gaattaacgg gynnnatccc tnttgggggg cnggnccccc cccctcgggg      480
ggttngggnc aggnccnaat tgtttaaggg tccgaaaaat cctcccnaga aaaaaancct      540
ccaggttgag nntnggggtt nccccccccc cangggccct ctognaaagt tggggttttg      600
ggggccctgg attttttttt ccttntttcc tccccccccc cngggganag aggttngagt      660
tttgnctnnc ggcocccnch aaganctttn ccgatttnan ttaaatccnt gcttngggga      720
agtcctttn agggntaaan ggcocccctm cggg

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<210> 21
<211> 755
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)... (755)
<223> n = A, T, C or G

<400> 21
atcancocat gaccccnac angggacncc tcanccggnc nnaacnccn ccggccnata      60
angtnagmnc actnccnttn naccancccc cncnccatd gcccncnanc cnaacgcncta      120
mcanatncc actganngcg ogangtnan ngagaaanct nataccanag nccocanacn      180
coagctgtcc nanhangcct nnnatccngg nnnatccaat ntgnancctc cnaagtattn      240
mncnccanat gattttccct anccgattac ccntnccccc tanccectcc cccccaacna      300
cgagggcnct ggncncaagg nngcncnccc ccgctagmtc cccnccaaat cncnccncta      360
aactcaneen nattacnccg ttctngayta tcactcccg aactcaocct taactaacct      420
aaaaancten gatcacaaat aatncaagcc tcnttatnac actntgactg ggtctctatt      480
ttagnggtcc ntnanccatc ctaatacctc cagttctnct tcncccaatt ccaanggtct      540
ctttngaca gaatnttttg gttcccnntt ggyttcttan ngaattgccc ttctnngaac      600
gggtctctct tttccttcgg ttancctggg ttcnncgggc cagttattat ttccnctttt      660
aaatctnnc cttttanttt tggcatttna aaccccgggc ctbgaaaagc gcccccgtgt      720
aaagggttgt tttganaaaa tttttgtttt gttccc

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<210> 22
<211> 849
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)... (849)

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<223> n = A,T,C or G

<400> 22

tttttttttt	tttttangtg	tngtngtgcg	ggtagagggt	taotacaant	gtgaanacgt	60
acgtngngan	taangcgacc	vganttttag	gannncocct	aasatcaaac	tgtgaagatn	120
atcctgmnna	cggaaggttc	acggngngat	natgotatggg	tgnccnctcc	cannnccttn	180
ataaatctng	nggcctcgcc	caccaccttc	ggcgccccng	ngnccggggcc	cgggtcattn	240
gnnttaacon	caetnngnca	ngggtttccn	ccccnnncng	acccngggca	tccgggggtnc	300
tctgtctctc	cctgnagncn	anasantggg	ccnccgncoc	ctttacccct	nnacaagcca	360
cnccctctta	noocnagccc	ccnctccact	nngggggact	gccannngct	cgtttnctng	420
nnacccnnnn	gggttccctg	gttgtogant	cnaccgnang	ccanggtatc	cnaagggaag	480
tgcgtttatg	gcccctaccc	ttcgtcnogg	nnccacccctc	ccgacnanga	noeqctnccg	540
cnvnnccngc	cctcnctctg	caacacccgc	actctnctgt	ncggnnnccc	ccccacccgc	600
acccctcncc	ngnccgnanc	ctccnccncc	gtctcannca	ccaccccccgc	cgcgcaggcc	660
ntcncacac	ggngnccngc	nagcncnctc	gcncccgcca	ggcncnccct	cgcncnngaa	720
ctactctngg	ccantnngcc	tcannccnna	cnaaacgcgc	ctgcggcgcc	cgnagcgccc	780
noctccncca	gtcctcccg	cttcnccccc	angntttccn	cgaggacacn	nnaccccgcg	840
nnccngcgg						845

<210> 23

<211> 872

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(872)

<223> n = A,T,C or G

<400> 23

ggcgaascta	taacttcgtc	gnactcgtgc	gcctcgtctc	ttttttcttc	cgnasaccatg	60
tctgaacanc	cagattnggc	ngatatcnan	aagntoganc	agtccaaact	gantaacaca	120
caacacnanc	agntasaatc	ncctgctctc	anagtanaen	attgaacnng	agaaocanpc	180
agggcgatcg	taatnaggcg	tgcgcgccca	atntgtcncc	gtttattatn	ccagctctnc	240
ctnccncccc	taactcttcn	nagctgtcnn	acccctngtn	cgnancccc	naggtcggga	300
tccgggtttan	antgscggng	cnnccctccc	ccnctccat	naagacccnc	cgcacacccc	360
nanngcnccg	ccccctnctc	cttcgcaccc	ctgtcctatn	ccctcgtcgc	ctggcnznng	420
accgcattga	ccctcgcnnn	cttncnngaaa	ncgnanaagc	cggggttggn	annanogctg	480
tgggnnccgg	tctgscacgc	gttccctccn	nnncttccca	ccactctctc	tacngsgtct	540
cnccgccttc	tcnnncaenc	ccctgggaagc	ttctctntgc	cccccttnac	tcccccctct	600
cgngctgccc	cgcacccacc	ntcatttnca	naagatcttc	acaaannccct	ggntnnctcc	660
cnanngcncc	gtccnccnag	ggagggggng	ggnnccnctg	nttgcagctg	ngggngngtc	720
cgaaantccc	tctnccctcan	cncctacccc	cggcggnnct	ctngtntncc	saacttancaa	780
ntctccctcg	nggcncatcc	tcagctctnc	ccncccccct	ctctgcantg	tnctctgtctc	840
tnacncttcc	gaattctcgn	cncctctctt	cc			872

<210> 24

<211> 815

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(815)

<223> n = A,T,C or G

<400> 24

gcctgcacgc	ttgagtattc	tatagngtca	ccaaatanc	ttggcctaat	catggctnta	60
------------	------------	------------	-----------	------------	------------	----

nctgnettec	tgigtcaaat	gtatacnaan	tanatatgaa	tctnatntga	caagamngta	120
tontncatta	gtacaantg	tnntgtccat	cctgtcngan	canatcccaa	tnnattnecn	180
cgatctcccn	gcnccatn	taatatgggaa	ntonnatnnn	ncacccnccat	ctatctntcc	240
gmcnccctgac	tggnagagat	ggatnnaatc	tnntatgacc	ncatgtgtca	tcttggtatn	300
aanaacccncc	cgcnccccac	cgyttngnng	cnagccnntc	ccaaagacctc	ctgtggaggt	360
aaacttggtc	agannatcaa	aaactgggaa	accgccnnc	angtnnaagt	ngnnncanan	420
gactccgtcc	aggnitnacc	atcccttenc	agcgccccc	ttngtgcctt	anagngnagc	480
gtgtccnanc	cactcaacat	ganacggccc	agmccanccg	caattingcca	caatgtccgc	540
gaaccccaata	gggggaatna	tncaaanccc	caggattgtc	cnccnangaa	atcccncaac	600
ccncccccac	ccncttttgg	gacnctgacc	aantcccgga	gtaccagtc	ggccnngctc	660
cccccacggg	naacctgggg	gggtgaanct	cngnatccac	cngnccaggn	ntcnaagga	720
acgggucctn	ggncgaanng	ancnntonga	agngccnct	cgtatacccc	cccctcncca	780
ccncaengnt	agntcccccc	cngggtncgg	aangg			815

<210> 25

<211> 775

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(775)

<223> n = A,T,C or G

<400> 25

ccgagatgct	tgctccgtg	gccttagctg	tgctccgct	actctctctt	tctggccctg	66
aggtctatcca	gggtactcca	aagattncgg	tttactcnog	tcactccgca	gagaatggaa	120
agtcnaattt	cctgaattgt	tatgtgtctg	ggtttccatc	atccgacatt	gaanttgact	180
tactgaagaa	tggnagagaa	attgaaaaag	tgagacatc	agccttgctc	tccagcaagg	240
actggtcttt	ctatctctng	tactacactg	aattccaccc	cactgaaaaa	gctgagtatg	300
ccgtgcgttg	gaacccatgt	actttgtcac	agccccagat	agtttaagtgy	gctcgagaca	360
tgtaaagcga	cnncatggaa	gtttgaagat	gacgcatttg	gattgggtga	attccaaatt	420
ctgctttgct	gcttttbaat	antgatatgc	ntataacccc	tacccttbat	gnocccaaat	480
tgtgggggtt	acatnangt	tcnctatnga	catgatcttc	ctttataant	cnccncttcg	540
aattgcccggt	cncccnngtn	ngaattgttc	cnnaaccacg	gttggctccc	ccagggtcncc	600
tcttaccggaa	gggcccgggc	cnctttncaa	ggttggggga	acnnaaaatt	tcnctnttgc	660
cnccncccca	cnntcttgng	nnccnacttt	ggaccccttc	cnattccctt	tggccnna	720
nccttnncta	anaaaacttn	aaanctngc	naaannttn	acttccccc	ttacc	775

<210> 26

<211> 820

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(820)

<223> n = A,T,C or G

<400> 26

anattatnac	agtgtaantc	tttccacag	gtgtgtanag	ggaaaggggc	ctagaggcat	60
ccanagata	nettatnaca	acaghtgctt	gaaccaagac	tgctggggac	atttctgca	120
gaaaaggtgy	cggtccccat	cactctctct	ctcccatagc	catccccagc	gggtgagtag	180
ccatcangcn	ttcggtggga	ggagctcang	gaacaacacn	accacagacc	anacagacca	240
ntgatgacca	tgggcggggg	ggagctctct	cctgtnaccc	gggtggcana	nganagccta	300
nctgaggggt	cacatataaa	acgttaacga	cnagatnanc	caactgcttc	aagtgcaccc	360
ttctacactg	acnacccggg	acnnnaact	gcngcctggg	gacgcctctg	gtcagaccta	420
acnnaagact	ccactgcccc	cccatggccc	taagntcccc	tggtctctgc	aagggaagct	480

cctctgttga	atnctgggga	naccaaagga	sccccctct	ccanctgtga	aggaaaaaann	540
gatggacttt	tccccctccg	gccnntcccc	tcttcttka	caagccacct	nttactatc	690
tccctctntt	ntactgncnc	acttttsacc	ccnnnatctt	ccttnattga	tgggaanctn	660
ganattccaa	tnacgcccnc	ctenatctng	naaanacnaa	sactatctna	ccccggggat	720
gggnaactcg	ntactctct	cttttctnct	acnccnntt	ctttgctct	ccttagatca	780
tccasccntc	gntggccctn	ccccccann	tcttttacc			820

<210> 27

<211> 818

<212> DNA

<213> Homo sapien

<220>

<221> misc feature

<222> (1)...(818)

<223> n = A,T,C or G

<400> 27

tctgggtgat	ggcctcttcc	tcttcaggga	cctctgactg	ctctggggca	aagaactct	60
tgtttcttct	ccagagcccca	gynagcgggtg	attcagccct	gcccaacctg	attctgatga	120
ctggggatgc	tgtgagggac	caaagggggca	aataggggtcc	caggggtccag	ggagggggcc	180
ctgctgagca	cttccgcccc	tcacctctgcc	cagccctctgc	catgagctct	ggcctgggttc	240
tcggctctca	gggttctgct	cttccangca	ngccanccaa	tgggctctgg	ccacactctg	300
ttctctctgc	ccctctctg	gctctganc	tctgtcttcc	tgtctctgct	angcnccttg	360
gatctcagtt	tccctctctc	anngaacctct	gtttctgann	tcttcannta	actntgantt	420
tatnacccan	tggnctgtnc	tgtcnnacttt	taattgggca	gacccgctaa	tcctctcctc	480
actcccttcc	anttcnana	acncccttnc	ctntctctcc	ccntancccg	ccnngggganc	540
ctccttctgc	ctnaccangg	gccnccnccg	ccctnctnct	ggggggggng	gttncctncc	600
ctgntncccc	cncfctcnnt	tnactctgtc	cncnccnccg	angcannctc	ngngtcccn	660
tnnctctctc	ngtnlognaa	ngctnctnct	tnnannngna	ngntnctnct	tcctctctcc	720
ccnctngang	tnnttannnc	ncnngncccc	nnnnccnnna	nggnantknn	tctnctnccg	780
ccccncccc	ngnattaagg	cctcnnctct	ccggccnc			818

<210> 28

<211> 731

<212> DNA

<213> Homo sapien

<220>

<221> misc feature

<222> (1)...(731)

<223> n = A,T,C or G

<400> 28

aggaagggcg	gagggatatt	gtanngggatt	gaggggatagg	agnataangg	gggaggtctg	60
tcocacacatg	anggtgnggt	tctcttttga	angaggggttg	ngtttttana	ccnggtgggt	120
gatfnasccc	catgtgatgt	agmnaaaggn	tttnagggat	tittctggctc	ttatcagtat	180
ntanattctc	gtnaatcgga	aatnatattt	tcnncnggaa	aatnttgcct	ccatcccgnaa	240
attntctccc	ggtagtgcct	nttngggggg	cngccangtt	tcccagggctg	ctanaactgt	300
actaaagntt	naagctggan	tnccaatgaa	acccctnccac	agagnatccn	taccagactg	360
tnncttncct	tcccccctng	actctgongn	agcccaatac	ccnngngnat	gtcccccngn	420
nnngcgncnc	tgaannnnnc	tcngngctnn	gancatcang	gggttctgca	tcaaaagcnn	480
cgtttcnact	naagggcactt	tngcctcctc	caacnctctg	ccctccncca	tttngccctc	540
nggttncct	acgctnnng	cncctnnntn	ganattttnc	cgcctngggg	naancctcct	600
gnaatgggta	gggncttntc	ttttnacenn	gggtntaact	aatcnctcnc	acgctactct	660
tctcnacccc	cccctttttt	caatccccc	ggcnaatggg	gtctcccnna	cngnnggggg	720
nnccccnnnc	c					731

<210> 29
 <211> 822
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)... (822)
 <223> n = A, T, C or G

<400> 29
 actagctcag tgtggtggaa ttccatfctg tigggggncc ttctatgcnt antnttagat 60
 cgtctcacc tcacancctc cccacnangc ctataangaa nannaataga notgtcnmt 120
 stntntacc tcabamcct cnnnacccac tccctcttaa cccctactgt gcctatngn 180
 tnnctantct ntgcgcctn cnanocccn gtgggcccac cncnagnatt ctcatctcc 240
 tcnccatntn gcttananta ngtncatacc ctataccctac nccaatgcta nnnctaanen 300
 tccctnattt annntaecta cccctgacct ngactttmc atnancctcc aatttgaatc 360
 tactctgact cccacngect annnatlaga anentccccc aacnatntct caacccaaac 420
 ntccacaccc talctancctg ttcccacccc nttnccctcc atcccnnac aacccccctc 480
 ccaaatcccc nccacrtgac nccatacccn caccatccc gcaagccnan gyncatttan 540
 cccctggant cccnctngga naaaaaaac cccnactctc tancnccnat ctccctaaana 600
 aatnctccn naatttactn nccntcccat caancccccn tgaacnnaa cccctgtttt 660
 tannctccct ctttgaana cccacccctt anncccccac ctttngggcc ccccccactnc 720
 ccccatgaag gcnccccaat cnangaaacg nccntgaaaa aacnagggcna anannntccg 780
 canatccctat cccctanttn ggggnccctt nccnngggcc cc 822

<210> 30
 <211> 787
 <212> DNA
 <213> Homo sapien
 <220>
 <221> misc_feature
 <222> (1)... (787)
 <223> n = A, T, C or G

<400> 30
 cggcgccctg ctctggcaca tgcctctga atggcatcaa aagtgatgga ctgcccattg 60
 ctgagagaag cctctctccc taactgcat atggagccct gcagactgag ggctcccctt 120
 gtctgagaga ttgatgtct gaagtctgtg agtgtggctt ggagctctcc atctacatna 180
 gctggagacc ctggagggcc tctctgcaca gccctcccct tctctccacg ctctccanng 240
 aacccagggg ctccaggaag cccattatc ccagnangac atggtgtttt tccacggcgg 300
 cccatggggc ctgnaagggc aggtctcct ttgacaccc ctctcccgtc ctgcccggga 360
 ggccgtggga tccactantt ctanaacggn ccccccncg gtggggagctc cagcttttgt 420
 tcccttaatt gaaggttaat tgnogcttg gctaatcat aggtcanaac tntttctctg 480
 gtgaaattgt tntccctc nonattccn nonactacn aaccoggaan cctaaagtgt 540
 taaagctggg ggttgcctn nngaatnaac taaactaat taattgcttt ggctcatggc 600
 cgtcttccn ttccggaaaa ctgtctccc ctgcttntat gaatgggcca ccccccnggg 660
 aaaaaggctt tgcnttttng gggntcctt cccctccccc cctcctnaan cccctnagct 720
 cgggtgttnc nggtngcggg gaanggggat nncctccnc aaagggggng agnangntat 780
 ccccaaa 787

<210> 31
 <211> 799
 <212> DNA
 <213> Homo sapien

<220>

<21> misc_feature
 <22> (1)...(799)
 <23> n = A,T,C or G

<400> 31
 tttttttttt tttttttggc gatgctaactg ttttaattgca ggaggtgggg gtgtgtgtac 60
 catgtatacag ggcattataga agcaagaagg aaggagggag ggcagagcgc cctgctgagc 120
 aacaaaggagc tccctgagcc ttctctgtct gtctcttggc gcaggcaact ggggaggcct 180
 ccgcaggggt gggggccacc agtccagggt tgggagcact acanaggggt ggaagtgggtg 240
 gtggtgtgtn cnaatggcct gncacnatic cctacgattc tggacacctg gatttcacca 300
 ggggacccctc tttctccca agynaaattc ntannatctn aaagaacaca actgtttctt 360
 cngnatttct ggtgtttcat ggaagacaca ggtgtccnat ttngctgggg actgtgtaca 420
 tatgtgtccg gcccctctct cccctcnaaa agtaattca ccccccccn cctctnttg 480
 cctggggccct taantaacca caccgggaact canttanta ttcattctng gatgggttg 540
 ntatnccncc cctgaagcgc ccaagttgaa aggcacagcc gtnccnctc cccatagnan 600
 ntttttncnt canctaatg ccccccgagc aacnatacaa tcccccccn tgggggcgcc 660
 agccccaggg ccccgctctg gnnnccngn cncgnantcc ccaggtcttc coantcngn 720
 cccnngcnc cccgacgcs gaacanaagg atngagcnc cgcannanan aggtncncc 780
 ctgcgcgcc cccnngnng 799

<210> 32
 <211> 789
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(789)
 <223> n = A,T,C or G

<400> 32
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 ggcacacagc tccggcgccg gcggggcggt cctcaactgc ggtcccaaat ntgcagcctc 180
 cgtcccgct tgaatnctct ctgcagctgc aggatgcctt aaaaacaggc ctcggcmtnt 240
 ngtgggccc ctgggatttn aatttccacg ggcacactgc ggtgcacnc cctcaccacc 300
 atagggaat agtggnttta cccnccnccg ttggcncact ccccttgaa accactnttc 360
 gcggctccgg catctgtct taacacttgc aaacnctggg gccctctttt tggttantnt 420
 nccngccaca atcatnaact agactggcnc gggctggccc caaaaaancn ccccaaaacc 480
 gghccatgic ttncgggggt tgcctgnatn tncatcact cccgggcnca ncaggncaac 540
 ccaaaagtctc ttggggcccn caaaaaancn cgggggggnc ccagtttcaa caaagtcact 600
 ccccttggcc ccccaactct ccccccgttt nctgggtttg ggaacccacg cctctncttt 660
 tggngccaaa gntggntccc ccttcgggpc cccggtgggc ccmctctaa ngaaaaancn 720
 ntcctncca ccatccccc nngnnaagnc tacaangns tccctttttt tanaaacggg 780
 ccccccng 789

<210> 33
 <211> 793
 <212> DNA
 <213> Homo sapien

<220>
 <221> misc_feature
 <222> (1)...(793)
 <223> n = A,T,C or G

<400> 33
 gacagaaact gttggatggt ggagacactt tctatacgc ttacaggaca gcagatgggg 60

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aatctaatggtg ttgttggaaga atanaaacccg agttctacga gctgctgctc aaaggacttg 120
gactaaagtc tgatgaactt cccaatcaga tgagcttgga tgaattggcca gaattgaana 180
agaagtttgc agatgtattt gcaaaagaaga cgaaggcaga grggtgtcaa atctttgagc 240
gcacagatgc ctgtgtgact cgggtttctga ctlttgagga gtttgttcat catgatcaca 300
acaagaacgc ggggtgtgtt atcaccantg aggagcagga cgtgagccoc cgccttgccac 360
ctctgtgttt aaacacccca gccatccctt ctctcaaaag ggtaccacta ctcttagagc 420
ggncgcaccc cgggttgagc tccagcttlt gttcccttba gtaggggtta attgpgcgt 480
tggcgtaate atggtcatan ctgtttctgc tgtgaattg ttatccgctc acaattccac 540
accacatagc acccggaagc atnaaatltt aaagccttgn ggtngcctaa tgaatgact 600
naactacatt aattggtctt gcctcactg cccgttttcc agtcgggaaa acctgtctct 660
gocagctgcc nttaaatgat ccggccaccc cccggggaaa aggcngtttg ctctttgggg 720
cgncctccc gotttctgc ttactgaant ccttcccccc ggtctttggg cttagggcna 780
acggtatcna cct 793

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<210> 34

<211> 756

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(756)

<223> n = A, T, C or G

<400> 34

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anacagtgcg ggaanaagct ggttgactc agctagtctt ttctggagct caactctttg 120
ccaaaccaag ggaaccaagc gaaccaaacg cagctaatct tggccagtg cactactggg 180
atcgggggcc aattggagct ctacgcgaan gacatccoot ccttcagagc ctacatgggc 240
cagctcaaat gctactactt gattacaaan gaggagctcc ccgagtcagc ctatatgcac 300
cagctcttgg gctcaaatct ccttctctg ctgtccnaga accgggtggc tgaantccac 360
acgganttgg ancggtgcgc tgcaccaaga cataccanac aatgtctaca tcnaccacca 420
gtgtcctgga gcaatactga tgganggcag ctaccncaaa gtnttctg ccnagggtaa 480
catcccccgc cgagagctac accttcttca ttgacatcct gcttcagact atcagggtg 540
aaatcgcneg ggttgctcca gaaggctcnc aanaaatcc ttttctctga aggcocccgg 600
atnctnagt nctagaatgc gcccgcatc ggggtgganc ctccaccctt tctgttacct 660
ttactgaggy ttatttgctg ccttggcgt tatcatggtc aacccngttn cctgtgttga 720
aattnttaac cccccaacat tccagccna catfng 756

```

<210> 35

<211> 834

<212> DNA

<213> Homo sapien

<220>

<221> misc_feature

<222> (1)...(834)

<223> n = A, T, C or G

<400> 35

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aacaggtatc tgcocctgaa gctctgggtg gctgtnttta agttgctcag tctccgctca 120
tgctcagaca cnetottggg caaaaaaan caggaattga gtcttgattt caactccaat 180
aatcttcnag gctgtctgct cgttgaactc gatgacnag gcacagctgt tgtgtntgat 240
aaattccanc angttctctt tggtagcttc cccctcaaa gttgtccgpc ctctcataaa 300
ctctnnaan angmananc canctttgtc gacttggnat ttggnaaca cgtcactgtt 360
ggaaactgat cccaaatggt agtcatcna tgcctctgct tgcctgcaaa aaactgtctt 420
ggcnaaactc cgaactcccn tcttgaag aagcnaatca cccccccctc cctggactcc 480

```

nncaangact	ctacocgetnc	ccntcccsng	cagggttngt	ggcansccgg	gcoontgggc	540
ttcttcagcc	agttccnhet	attcatcagc	cactctgcga	gcgtttntai	tccttggggg	600
ggaanccgic	tcctccctcc	tgaannaact	ttgacccgtag	gaatagccgc	gcntcncct	660
acninctggg	cagggttcaa	antccctccn	ttgncnntan	cctcggggcc	ttctgggatt	720
noonaacttt	ttctctcccc	cnccccnngg	ngtttggntt	ttccannggy	cccccactct	780
gctattggcc	antccctggg	gggcntntan	cacccactnt	ggtcccttag	ggcc	834

<210> 36

<211> 814

<212> DNA

<213> Homo sapien

<220>

<221> misc feature

<222> (1)...(814)

<223> n = A,T,C or G

<400> 36

cggnccgttt	ccngccgggc	cccgtttcca	tgaacnaagcc	tcoccttcang	ttaaatacnn	60
ccagaaaac	attcaatggg	tgctctacta	atcacatnsta	cnaaccagta	agcctggcca	120
naacccgcaac	ctaggccact	cctacccaag	gaagaaagcc	tggtctctcc	acccocctcta	180
ggaaagggct	gccttgttag	acaccacaat	ncggctgaat	ctnaagctctt	gtgttttact	240
aatgaaaaaa	aaaaataaac	aaanaggttt	gtctctctgg	ctgcccacog	cagcctggca	300
ctaaaaacano	ccagcgctca	ctctcgcttg	ganaaatatt	ctttgctctt	ttggacatga	360
gggttgatgg	tatacaatgoc	acntttccac	ccagctggggc	acccctcccc	catttttgtc	420
antgancctgg	aaggccctgaa	ncttagcttc	caaaagtctc	ngcccacacg	acgggcccac	480
aggggagctc	ntttncagtg	gatctgcgaa	aanatacccn	tatcatcnn	gaataaaaag	540
gcccctgaac	ganatgcttc	cancancctt	taagacccat	aatcctngaa	ccatgggtgc	600
cttcgggttc	gatccnaaag	gaatgttact	gggtcccnat	cctccttttg	ttncctacgt	660
tgntttggac	ccntgcingn	atncccaam	tgsnatcccc	agaagcaccc	ttccctggc	720
atttganttt	cntaaattct	ctgcctacn	ncgaaagca	cnattccctn	ggccncaan	780
ggngaacctc	agaaggtctn	ngaaaaacca	cncn			814

<210> 37

<211> 760

<212> DNA

<213> Homo sapien

<220>

<221> misc feature

<222> (1)...(760)

<223> n = A,T,C or G

<400> 37

gcattgctgc	cttccctcaa	gttgttcttg	ttgcataac	aaccacacta	ggtaaagggy	60
gcgcagtggt	cgctgaaggy	gttgtagtac	cagcgcggga	tgctctcctt	gcagagtctc	120
gtgctctggca	ggtccacgca	atgcoccttg	tcactgggga	aatggatgdy	ctggagctog	180
tcaaacccga	tcgtgtattt	ttcacangca	gcctctccgg	aagctccggg	gcagltgggg	240
gtgtgtctac	actccactaa	actgtcgatn	cacnagccca	ttgctgcagc	ggaactgggt	300
gggctgacag	gtgcacgaac	acactggata	ggcctttcca	tggaaagggc	tgggggaat	360
cncctnancn	caaaactgct	ctcgaaggcc	accttgacac	cccgcacagg	ctagaatgc	420
actcttcttc	ccaaagtgat	ttgtttttgt	tgcccacgca	ncctccanca	aaccaaaanc	480
ttgcaaaatc	tgctccgtgg	ggttcctnnn	taocangytt	ggggaaaana	acccggcmgn	540
gancnccttt	gtttgaatgc	naaggnaata	atcctcctgt	cttgcttggg	tggaaanaga	600
caattgaact	gttaacnttg	ggccgggttc	cnotnggggt	gtctgaact	aatcccgctc	660
acttgaaaaa	ggtangtgc	ttccttgaat	tcocaaantt	ccctngnttt	tggttntatt	720
ctcctctncc	ctaaaaatog	ttttccccc	cctnanggg			760


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<210> 38
<211> 724
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(724)
<223> n = A,T,C or G

<400> 38
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cttcocnaaat tgtocnaaoc ootcnmocaa stnnccattt cggggggggg gtcccaaac 120
caaatataat ttggaaattta aattaaatnt tnatnggggy aanaaancaa atgtnaagaa 180
aatitaaocm attatnaact taatatnoctn gaacccocntg gnttccaaaa sttttlaaoc 240
otaaatoccc tccgaaatgy ntaanggaan accaaattcn cctaaggctn tttgaaggtt 300
ngatttaaac cccctttaat tnttttnacc cnagnctnaa ntatfngnt tccggtgttt 360
tccnttaan cntnggtaac tcccgntaat gaannccot aanncaatta aaccgaattt 420
ttttgaaatt ggaasttccn nggggaattna cgggggtttt tcccttttgg ggggcatncc 480
cccnotttgg ggytttgggo ntagggttga tttttmang ncccaaaaaa nccccanaa 540
aaaaaatccc caagnattaa ttngaatncc ccccttcccc ggccttttgg gaaggngggg 600
tttttggggg ccngganntt cnttccoccn ttncncccc ccccccnggt aaanggttat 660
agnntttggt ttttggggcc cttnanggac ottccggatn gaaattaaat ccccgggngc 720
gcgc 724

<210> 39
<211> 751
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(751)
<223> n = A,T,C or G

<400> 39
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caacacataa ttattttcaat ttgttttttt tatttttttt tttttttttt ctgctgtgtg 120
tttttttttt tttaactgaaa gtgagaggga acttttttgg cctttttttc tttttctgta 180
ggccggctta agottttctaa atttggaaaca tctaaagcaag ctgaanggaa aagggygttt 240
cgcaaaatga ctggggggaa aggaaggytt gotttgttaa tcatgaccta ttcgtgggtg 300
ttaactgctt gtacaaattac nttaacattt taattaattg tgcnaaango ttttaattana 360
cttgggygtt ccttccocan acccaacccn ctgacaaaaa gtgcngnccc tcaaatnatg 420
tcccgccant cnttgaacaa cccngungaa ngttctcatt ctcccncccc caggtnaaaa 480
tgaaaggyta caatntttas cncacacccc cnttggcnna gctgaatcc tcnaaaancc 540
ccctcaanen aakttnctng ccccggttnc gentangtcc cncccgggct cggggaantn 600
caccccngaa annctntnnc naccnaaatt ccgaaatat tcccnctcnc tcaattcccc 660
cncagaatnt cctcnncnan cncaattttc ttttntcacc gaacnngnnc cnnaaaatgn 720
annnccctc cncntngtccn naatcnccan c 751

<210> 40
<211> 753
<212> DNA
<213> Homo sapien

<220>
<221> misc_feature
<222> (1)...(753)

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